

## SEARCH REQUEST FORM

Requestor's Name: \_\_\_\_\_ Serial Number: \_\_\_\_\_  
Date: \_\_\_\_\_ Phone: \_\_\_\_\_ Art Unit: \_\_\_\_\_

**Search Topic:**

Please write a detailed statement of search topic. Describe specifically as possible the subject matter to be searched. Define any terms that may have a special meaning. Give examples or relevant citations, authors keywords, etc., if known. For sequences, please attach a copy of the sequence. You may include a copy of the broadest and/or most relevant claim(s).

**STAFF USE ONLY**

Date completed: <u>02-14-03</u>	Search Site	Vendors
Searcher: <u>Beverly e4994</u>	<u>STIC</u>	<u>IG Suite</u>
Terminal time: <u>22</u>	<u>CM-1</u>	<u>STO</u>
Elapsed time: <u></u>	<u>Pre-S</u>	<u>Dialog</u>
CPU time: <u></u>	<u></u>	<u>APS</u>
Total time: <u>25</u>	<u>N.A. Sequence</u>	<u>Geninfo</u>
Number of Searches: <u></u>	<u>A.A. Sequence</u>	<u>SDC</u>
Number of Databases: <u>1</u>	<u>Structure</u>	<u>DARC/Questel</u>
	<u>Bibliographic</u>	<u>Other CGN</u>

Jiang, D.  
091756690

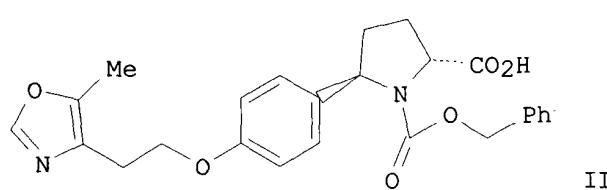
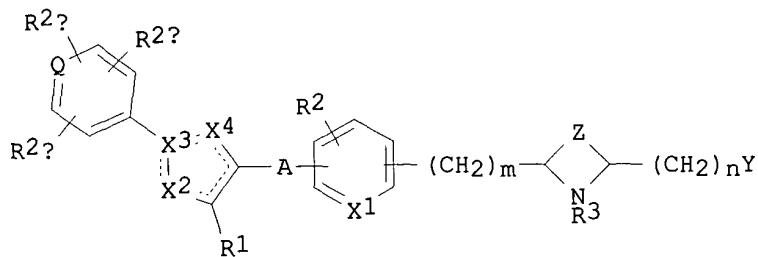
09/756690

FILE 'REGISTRY' ENTERED AT 10:31:04 ON 14 FEB 2003  
L1 31 SEA ABB=ON PLU=ON HGEGTFTSDLSKQMEEAVRLFIEWLKNGGPSSGAPP  
PS/SQSP

FILE 'HCAPLUS' ENTERED AT 10:31:51 ON 14 FEB 2003  
L2 40 SEA ABB=ON PLU=ON L1

L2 ANSWER 1 OF 40 HCAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 2002:927184 HCAPLUS  
DOCUMENT NUMBER: 138:14048  
TITLE: Preparation of oxazolylethoxyphenylprolines and  
related compounds as antidiabetic and  
antiobesity agents.  
INVENTOR(S): Cheng, Peter T.; Jeon, Yoon; Wang, Wei  
PATENT ASSIGNEE(S): Bristol-Myers Squibb Company, USA  
SOURCE: PCT Int. Appl., 107 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002096357	A2	20021205	WO 2002-US16628	20020523
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			
PRIORITY APPLN. INFO.:			US 2001-294505P	P 20010530
OTHER SOURCE(S):		MARPAT 138:14048		
GI				



AB Title compds. [I; m, n = 0-2; Q = C, N; A =  $(CH_2)_x$ ,  $(CH_2)_{x1}$ , with an alkenyl or alkynyl bond in the chain,  $(CH_2)_{x2}(CH_2)_{x3}$ ; x = 1-5; x1 = 2-5; x2, x3 = 0-5; provided that  $x_2 \geq 1$  of x2 and x3  $\neq 0$ ; X1 = CH, N; X2 = C, N, O, S; X3 = C, N; X4 = C, N, O, S provided that  $x_1 \geq 1$  of X2, X3, X4 = N; in each of X1-X4, C may include CH; R1 = H, alkyl; R2 = H, alkyl, alkoxy, halo, (substituted) amino; R2a, R2b, R2c = H, alkyl, alkoxy, halo, (substituted) amino; R3 = H, alkyl, arylalkyl, aryloxycarbonyl, alkyloxycarbonyl, alkynylloxycarbonyl, alkenyloxycarbonyl, arylcarbonyl, alkylcarbonyl, aryl, heteroaryl, cycloheteroalkyl, heteroarylcarbonyl, heteroarylhetereoarylalkyl, alkylcarbonylamino, arylcarbonylamino, heteroarylcarbonylamino, alkoxy carbonylamino, aryloxycarbonylamino, heteroaryloxycarbonylamino, heteroarylhetereoarylcarbonyl, alkylsulfonyl, alkenylsulfonyl, heteroaryloxycarbonyl, cycloheteroalkyloxycarbonyl, aryloxylhetereoarylalkyl, heteroarylalkyloxylarylkyl, arylarylkyl, arylalkenylarylkyl, arylaminoarylalkyl, etc.; Y = CO2R4, 1-tetrazolyl, P(O)(OR4a)R5, P(O)(OR4a)2; R4 = H, alkyl, prodrug ester; R4a = H, prodrug ester; R5 = alkyl, aryl; Z =  $(CH_2)_{x4}$ ,  $(CH_2)_{x5}$ ,  $(CH_2)_{x6}(CH_2)_{x7}$ ; x4 = 1-5; x5 = 2-5; x6, x7 = 0-4], were prepd. as antidiabetic and antiobesity agents (no data). Thus, title compd. (II) was prepd. in 6 steps.

IT 141758-74-9, AC 2993

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (coadministration; prepn. of oxazolylethoxyphenylprolines and related compds. as antidiabetic and antiobesity agents)

L2 ANSWER 2 OF 40 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2002:832649 HCAPLUS

DOCUMENT NUMBER: 137:346934

TITLE: Methods for treating conditions associated with insulin resistance by administering a GLP-1 compound

INVENTOR(S): Holst, Jens Juul; Olsen, Mette Zander; Hathaway, David R.

PATENT ASSIGNEE(S): Restoragen, Inc., USA

SOURCE: PCT Int. Appl., 60 pp.

CODEN: PIXXD2

09/756690

DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002085406	A1	20021031	WO 2002-US13088	20020424
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			

PRIORITY APPLN. INFO.: US 2001-285699P P 20010424

AB The present invention relates to methods and compns. for treating insulin-assocd. conditions comprising administering a glucagon-like peptide-1 (GLP-1) compd. to subjects suffering therefrom. The insulin resistance-assocd. condition of the invention is type-2 pre-diabetes, atherosclerotic cardiovascular disease, drug-induced insulin resistance, congestive heart failure, diminished exercise capacity of skeletal muscle, and left ventricular dysfunction with cardiac metabolic myopathy or diminished exercise capacity of skeletal muscle; with the proviso that said congestive heart failure is not assocd. with toxic hypervolemia.

IT 474444-81-0

RL: PRP (Properties)

(unclaimed protein sequence; methods for treating conditions assocd. with insulin resistance by administering a GLP-1 compd.)

REFERENCE COUNT: 11 THERE ARE 11 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 3 OF 40 HCAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 2002:813924 HCAPLUS  
DOCUMENT NUMBER: 137:311200  
TITLE: Preparation of 2,1-oxazoline and 1,2-pyrazoline-based inhibitors of dipeptidyl peptidase IV  
INVENTOR(S): Sulsky, Richard B.; Robl, Jeffrey A.  
PATENT ASSIGNEE(S): Bristol-Myers Squibb Company, USA  
SOURCE: PCT Int. Appl., 61 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002083128	A1	20021024	WO 2002-US10936	20020405
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD,			

Searcher : Shears 308-4994

09/756690

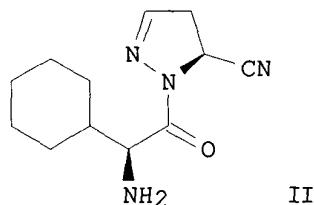
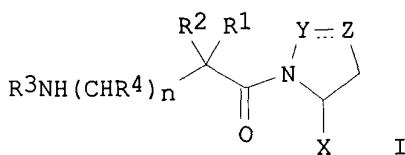
GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ,  
LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ,  
NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ,  
TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM,  
AZ, BY, KG, KZ, MD, RU, TJ, TM  
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE,  
CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT,  
SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE,  
SN, TD, TG

US 2002183367 A1 20021205 US 2002-107279 20020326

PRIORITY APPLN. INFO.: US 2001-283438P P 20010412

OTHER SOURCE(S): MARPAT 137:311200

GI



AB The invention describes dipeptidyl peptidase IV (DP 4) inhibiting compds. I [n is 0 or 1; X is H or CN; Y is N, NH or O; Z is CH2 when Y is O or NH, with Y-Z forming a single bond, and Z is CH when Y is N, with Y-Z forming a double bond; R1-R4 = H, alkyl, alkenyl, alkynyl, cycloalkyl, cycloalkylalkyl, bicycloalkyl, bicycloalkylalkyl, alkylthioalkyl, arylalkylthioalkyl, cycloalkenyl, aryl, aralkyl, heteroaryl, heteroarylalkyl, cycloheteroalkyl or cycloheteroalkylalkyl, which may be substituted; R1 may combine with R3 or R4 to form a ring (CR5R6)2-6 or (CR7R8)3-6, resp., where R5-R8 = H, OH, alkoxy, alkyl, aryl, etc.] and their pharmaceutically-acceptable salts or prodrug esters. A method is also provided for treating diabetes and related diseases, employing a DP 4 inhibitor I, optionally in combination with other therapeutic agents, including an antidiabetic, hypolipidemic, or anti-obesity agent. Thus, coupling of sultam-protected 1,2-pyrazoline-3-carboxamide with (S)-N-(tert-butoxycarbonyl)cyclohexylglycine (HOAt, Et3N, and EDAC in CH2Cl2), followed by sultam cleavage with methanolic ammonia, amide conversion to nitrile using imidazole, and deprotection, afforded II.TFA.

IT 141758-74-9, AC2993

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(antidiabetic agent; prepn. of oxazoline and pyrazoline-based  
inhibitors of dipeptidyl peptidase IV)

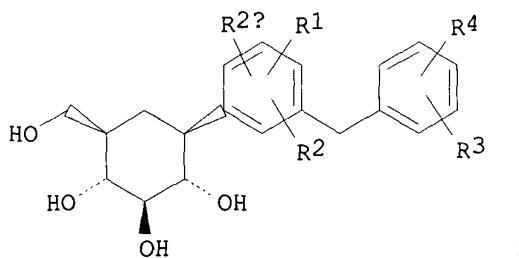
09/756690

REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 4 OF 40 HCAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 2002:813874 HCAPLUS  
DOCUMENT NUMBER: 137:311199  
TITLE: Amino acid complexes of C-aryl glucosides for treatment of diabetes  
INVENTOR(S): Gougoutas, Jack Z.  
PATENT ASSIGNEE(S): Bristol-Myers Squibb Company, USA  
SOURCE: PCT Int. Appl., 80 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002083066	A2	20021024	WO 2002-US11066	20020408
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				

PRIORITY APPLN. INFO.: US 2001-283097P P 20010411  
OTHER SOURCE(S): MARPAT 137:311199  
GI



AB Cryst. complexes are obtained from 1:1 or 2:1 mixts. of either the (D) or (L) enantiomer of natural amino acids and compds. of formula I [R1, R2, R2a = H, OH, OR5, alkyl, OCHF2, OCF3, SR5a, halogen; R3, R4 = H, OH, OR5b, alkyl, cycloalkyl, CF3, OCHF2, OCF3, halogen, CONR6R6a, CO2R5c, CO2H, COR6b, CH(OH)R6c, CH(OR5d)R6d, CN, NHCOR5e, NHSO2R5f, NHSO2-aryl, SR5g, SOR5h, SO2R5i, or a five, six or seven membered heterocycle which may contain 1 to 4 heteroatoms (N, O, S, SO, and/or SO2), or R3 and R4 together with the carbons to which they are attached form an annelated five, six or seven membered

09/756690

carbocycle or heterocycle which may contain 1 to 4 heteroatoms in the ring; R5, R5a-R5i are independently alkyl; R6, R6a-R6d are independently H, alkyl, aryl, alkylaryl or cycloalkyl, or NR6R6a form an annelated five, six or seven membered heterocycle which may contain 1 to 4 heteroatoms in the ring]. A method is also provided for treating diabetes and related diseases employing an SGLT2 (sodium dependent glucose transporters found in the intestine and kidney) inhibiting amt. of the above complex alone or in combination with another antidiabetic agent or other therapeutic agent. Thus, I (R1 = 4-Me, R4 = 4-OCHF2, R2, R2a, R3 = H) was prep'd. by a multistep procedure starting from o-toluic acid, anisole, 2,3,4,6-tetra-O-benzyl-.beta.-D-glucolactone, and CHF2Cl and treated with L-phenylalanine to form the cryst. 1:1 complex.

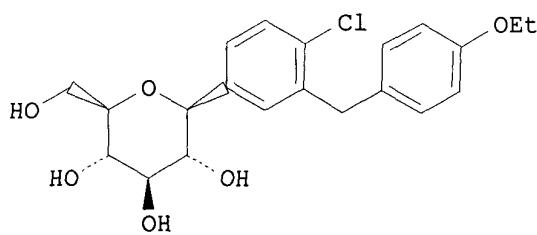
IT 141758-74-9, AC2993

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (prepn. of amino acid/C-aryl glucoside complexes for treatment of diabetes and related diseases)

L2 ANSWER 5 OF 40 HCAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 2002:736927 HCAPLUS  
DOCUMENT NUMBER: 137:247879  
TITLE: Preparation of antidiabetic agents C-aryl glucoside as human SGLT2 inhibitors  
INVENTOR(S): Ellsworth, Bruce; Washburn, William N.; Sher, Philip M.; Wu, Gang; Meng, Wei  
PATENT ASSIGNEE(S): USA  
SOURCE: U.S. Pat. Appl. Publ., 17 pp., Cont.-in-part of U.S. 6,414,126.  
CODEN: USXXCO  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 2  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2002137903	A1	20020926	US 2002-151436	20020520
US 6515117	B2	20030204		
US 6414126	B1	20020702	US 2000-679027	20001004
PRIORITY APPLN. INFO.:			US 1999-158773P	P 19991012
			US 2000-194615P	P 20000405
			US 2000-679027	A2 20001004

GI



AB An SGLT2 inhibiting compd. is provided having the formula I method is also provided for treating diabetes and related diseases employing an SGLT2 inhibiting amt. of the above compd. alone or in combination with another antidiabetic agent or other therapeutic agent (no data). 1A pharmaceutical combination comprising an SGLT2 inhibitor compd. and an antidiabetic agent other than an SGLT2 inhibitor, for treating the complications of diabetes, an anti-obesity agent, an antihypertensive agent, an antiplatelet agent, an antiatherosclerotic agent, and/or a lipid-lowering agent (no data). A method for treating or delaying the progression or onset of diabetes, diabetic retinopathy, diabetic neuropathy, diabetic nephropathy, delayed wound healing, insulin resistance, hyperglycemia, hyperinsulinemia, elevated blood levels of fatty acids or glycerol, hyperlipidemia, obesity, hypertriglyceridemia, Syndrome X, diabetic complications, atherosclerosis or hypertension, or for increasing high d. lipoprotein levels, which comprises administering to a mammalian species in need of treatment a therapeutically effective amt. of a compd (no data).

IT 141758-74-9, AC2993

RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(prepn. of antidiabetic agents C-aryl glucosides as human SGLT2 inhibitors)

L2 ANSWER 6 OF 40 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2002:640378 HCAPLUS

DOCUMENT NUMBER: 137:346599

TITLE: Cellular specificity of proexendin-4 processing in mammalian cells in vitro and in vivo

AUTHOR(S): Adatia, F. A.; Baggio, L. L.; Xiao, Q.; Drucker, D. J.; Brubaker, P. L.

CORPORATE SOURCE: Department of Physiology, University of Toronto, Toronto, ON, M5S 1A8, Can.

SOURCE: Endocrinology (2002), 143(9), 3464-3471  
CODEN: ENDOAO; ISSN: 0013-7227

PUBLISHER: Endocrine Society

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Glucagon-like peptide-1 (GLP-1) is a potent stimulator of glucose-dependent insulin secretion. Exendin-41-39 (Ex-4), isolated from Gila monster venom, is a highly specific GLP-1 receptor agonist that exhibits a prolonged duration of action in vivo. Although the processing mechanisms underlying liberation of GLP-1 from its prohormone have been elucidated, those for Ex-4 remain unknown. To examine the requirements for proEx-4 processing in mammalian cells, BHK fibroblasts, InR1-G9 islet A cells, and AtT-20 corticotrophs, which express different prohormone convertases (furin, prohormone convertase 2, and prohormone convertase 1, resp.) were transfected with full-length lizard proEx-4, and the processing of proexendin was examd. by HPLC and RIA. All of the transfected cell lines exhibited Ex-4-like immunoreactivity in the media, and Ex-4-like immunoreactivity was detected in exts. of InR1-G9 and AtT-20 cells. However, only media and exts. from AtT-20 cells (not InR1-G9 and BHK cells) contained a single peak by HPLC corresponding to synthetic Ex-4. To establish whether proEx-4 can be processed to Ex-4 in nonimmortalized mammalian cells in vivo, the mol. forms of exendin-4 were examd. in male and female mice expressing a metallothionein-proEx-4 transgene. ProEx4 mRNA transcripts were

09/756690

detected by RT-PCR in a broad range of both endocrine and nonendocrine tissues. Ex-4-like immunoreactivity was detected in pituitary, fat, adrenals, and testes; however HPLC analyses demonstrated that processed Ex-4 was found only in adrenals and testes. These results indicate that lizard proEx-4 is processed to mature bioactive Ex-4 in both rodent endocrine and non-endocrine mammalian cell types in vitro and in murine tissues in vivo. These findings may be useful for engineering cells that express a lizard proEx-4 transgene for the treatment of type 2 diabetes.

IT 188265-76-1, Exendin 4, pro- (Heloderma suspectum)  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(proexendin-4 processing to mature bioactive exendin-4 in rodent endocrine and non-endocrine mammalian cell types in vitro and in murine tissues in vivo)  
REFERENCE COUNT: 47 THERE ARE 47 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 7 OF 40 HCAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 2002:540258 HCAPLUS  
DOCUMENT NUMBER: 137:109267  
TITLE: Preparation of benzoxepinopyridines as HMG-CoA reductase inhibitors  
INVENTOR(S): Robl, Jeffrey A.; Chen, Bang-chi; Sun, Chong-qing  
PATENT ASSIGNEE(S): USA  
SOURCE: U.S. Pat. Appl. Publ., 42 pp., Cont.-in-part of U.S. Ser. No. 875,155.  
CODEN: USXXCO  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 2  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2002094977	A1	20020718	US 2001-7407	20011204
US 2002013334	A1	20020131	US 2001-875155	20010606
PRIORITY APPLN. INFO.:			US 2000-211595P	P 20000615
			US 2001-875155	A2 20010606
OTHER SOURCE(S):		MARPAT 137:109267		
GI				

\* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT \*

AB Title compds. I [X = O, S, SO, SO<sub>2</sub>, NR<sub>7</sub>; Z = HOCHCH<sub>2</sub>CH(OH)CH<sub>2</sub>CO<sub>2</sub>R<sub>3</sub>, 4-hydroxy-2-oxopyran-6-yl, etc.; n = 0, 1; R<sub>1</sub>, R<sub>2</sub> = alkyl, arylalkyl, cycloalkyl, alkenyl, cycloalkenyl, aryl, heteroaryl, cycloheteroalkyl; R<sub>3</sub> = H, alkyl, metal ion; R<sub>4</sub> = H, halo, CF<sub>3</sub>, etc.; R<sub>7</sub> = H, alkyl, aryl, alkanoyl, aroyl, alkoxy carbonyl, etc.; R<sub>9</sub>, R<sub>10</sub> = H, alkyl], were prep'd. as HMG CoA reductase inhibitors active in inhibiting cholesterol biosynthesis, modulating blood serum lipids such as lowering LDL cholesterol and/or increasing HDL cholesterol, and treating hyperlipidemia, hypercholesterolemia, hypertriglyceridemia and atherosclerosis (no data). E.g., a

IT multistep synthesis of II is reported.  
**141758-74-9**, AC2993  
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (coadministered agents; prepn. of benzoxepinopyridines as HMG-CoA reductase inhibitors for the treatment of hyperlipidemia, hypercholesterolemia, hypertriglyceridemia, atherosclerosis, and other disorders)

L2 ANSWER 8 OF 40 HCPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 2002:449715 HCPLUS  
 DOCUMENT NUMBER: 137:28591  
 TITLE: Preparation of GLP-1 fusion proteins for use in treating diabetes mellitus and other conditions  
 INVENTOR(S): Glaesner, Wolfgang; Micanovic, Radmilla; Tschang, Sheng-Hung Rainbow  
 PATENT ASSIGNEE(S): Eli Lilly and Company, USA  
 SOURCE: PCT Int. Appl., 200 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002046227	A2	20020613	WO 2001-US43165	20011129
W: AE, AG, AL, AM, AT, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, CZ, DE, DK, DK, DM, DZ, EC, EE, EE, ES, FI, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
AU 2002026897	A5	20020618	AU 2002-26897	20011129
PRIORITY APPLN. INFO.:			US 2000-251954P	P 20001207
			WO 2001-US43165	W 20011129

OTHER SOURCE(S): MARPAT 137:28591  
 AB The present invention relates to glucagon-like peptide-1 compds. fused to proteins that have the effect of extending the in vivo half-life of the peptides. The heterologous fusion proteins of the invention comprise a GLP-1 compd. fused to human albumin, a human albumin analog or fragment, the Fc portion of an Ig, or an analog or fragment of the Fc portion of an Ig. These fusion proteins can be used to treat non-insulin dependent diabetes mellitus as well as a variety of other conditions. Pharmaceutical formulations contg. the fusion proteins and polynucleotides encoding the proteins are also claimed.

IT **437124-38-4P 437124-39-5P 437124-51-1P**  
**437124-53-3P**  
 RL: BPN (Biosynthetic preparation); PAC (Pharmacological activity); PKT (Pharmacokinetics); PRP (Properties); PUR (Purification or recovery); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

09/756690

(amino acid sequence; prepn. of GLP-1 fusion proteins for use in treating diabetes mellitus and other conditions)

IT 435950-95-1DP, fusion proteins contg.

RL: BPN (Biosynthetic preparation); PAC (Pharmacological activity); PKT (Pharmacokinetics); PRP (Properties); PUR (Purification or recovery); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of GLP-1 fusion proteins for use in treating diabetes mellitus and other conditions)

L2 ANSWER 9 OF 40 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2002:392237 HCAPLUS

DOCUMENT NUMBER: 136:401651

TITLE: Preparation of fused pyridine derivatives as HMG-CoA reductase inhibitors

INVENTOR(S): Robl, Jeffrey A.; Chen, Bang-Chi; Sun, Chong-Qing

PATENT ASSIGNEE(S): USA

SOURCE: U.S. Pat. Appl. Publ., 46 pp., Cont.-in-part of U.S. Ser. No. 875,218.

CODEN: USXXCO

DOCUMENT TYPE: Patent

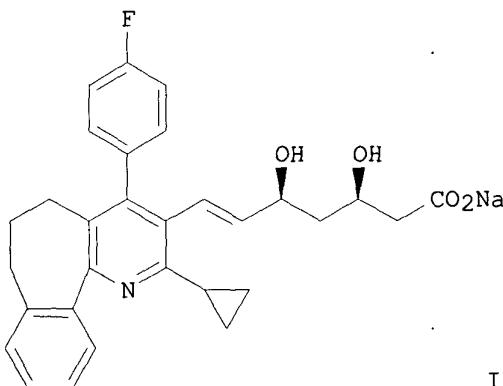
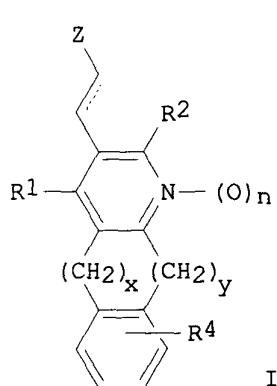
LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2002061901	A1	20020523	US 2001-8154	20011204
US 2002028826	A1	20020307	US 2001-875218	20010606
PRIORITY APPLN. INFO.:			US 2000-211594P P	20000615
			US 2001-875218	A2 20010606

OTHER SOURCE(S): MARPAT 136:401651  
GI



AB The title compds. I and their pharmaceutically acceptable salts, esters, prodrug esters, and stereoisomers are claimed [wherein: Z =  $\text{CH}(\text{OH})\text{CH}_2\text{CR}(\text{OH})\text{CH}_2\text{CO}_2\text{R}_3$  or corresponding pyranone lactone derivs.; n = 0, 1; x = 0, 1, 2, 3, or 4; y = 0, 1, 2, 3 or 4, provided that at least one of x and y is other than 0; and optionally one or more

carbons of  $(CH_2)^x$  and/or  $(CH_2)^y$  together with addnl. carbons form a 3 to 7 membered spirocyclic ring; R1, R2 = alkyl, arylalkyl, cycloalkyl, alkenyl, cycloalkenyl, aryl, heteroaryl, cycloheteroalkyl; R3 = H or lower alkyl; R4 = H, halo, CF<sub>3</sub>, OH, alkyl, alkoxy, CO<sub>2</sub>H, (un)substituted NH<sub>2</sub>, cyano, (un)substituted CONH<sub>2</sub>, etc.; R7 = H, alkyl]. The compds. are HMG-CoA reductase inhibitors, and are active in inhibiting cholesterol biosynthesis and modulating blood serum lipids, for example, lowering LDL cholesterol and/or increasing HDL cholesterol (no data). They are thus useful in treating hyperlipidemia and dyslipidemia, in hormone replacement therapy, and in treating hypercholesterolemia, hypertriglyceridemia and atherosclerosis, as well as Alzheimer's disease and osteoporosis. Preps. of several compds. are described. For instance, a multistep synthesis of fused pyridine deriv. II is reported. Compds. I may be used in a manner similar to atorvastatin, pravastatin, simvastatin, etc. Combinations of compds. I with various other drugs are claimed, the latter being specified as certain pharmacol. classes, as inhibitors of specific enzymes, as (ant)agonists of specific receptors, and as numerous named drugs.

IT 141758-74-9, AC2993

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (therapeutic compns. also contg.; prepn. of fused pyridine derivs. as HMG-CoA reductase inhibitors)

L2 ANSWER 10 OF 40 HCPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 2002:361520 HCPLUS  
 DOCUMENT NUMBER: 137:88643  
 TITLE: Endoproteolysis by isolated membrane peptidases reveal metabolic stability of glucagon-like peptide-1 analogs, exendins-3 and -4  
 AUTHOR(S): Thum, A.; Hupe-Sodmann, K.; Goke, R.; Voigt, K.; Goke, B.; McGregor, G. P.  
 CORPORATE SOURCE: Institute of Physiology, Philipps-University, Marburg, D-35037, Germany  
 SOURCE: Experimental and Clinical Endocrinology & Diabetes (2002), 110(3), 113-118  
 CODEN: ECEDFQ; ISSN: 0947-7349  
 PUBLISHER: Johann Ambrosius Barth  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 AB These in vitro studies aimed to characterize the pattern and the kinetics of endoproteolysis of the insulinotropic hormone glucagon-like peptide-1 (GLP-1) and related peptides by native ectopeptidases. Peptides were incubated with isolated rat or pig kidney brush-border microvilli membranes, which are a rich source of the ectopeptidases that are responsible for the post-secretory metab. of peptide hormones. The proteolytic products were sep'd. by reversed-phase HPLC column chromatog. and characterized by mol. mass and primary structure. The relative importance of specific peptidases was established by measuring the effects of specific peptidase inhibitors on the kinetics of proteolysis. Dipeptidyl-peptidase-IV was found to be rate-limiting in the endoproteolysis of GLP-1. GLP-1 homologs, exendins-3 and -4, exhibited exceptional stability in the presence of isolated kidney microvilli membranes. Our finding that exendin-4 is several orders of magnitude more stable than GLP-1 and Ser-8-GLP-1 is esp. noteworthy given this peptide's widely reported insulinotropic

09/756690

potency.

IT **141758-74-9**, Exendin 4 (*Heloderma suspectum*)  
RL: BSU (Biological study, unclassified); PKT (Pharmacokinetics);  
BIOL (Biological study)  
(endoproteolysis by isolated membrane peptidases reveal metabolic  
stability of glucagon-like peptide-1 analogs and exendin-3 and  
-4)

REFERENCE COUNT: 32 THERE ARE 32 CITED REFERENCES AVAILABLE  
FOR THIS RECORD. ALL CITATIONS AVAILABLE  
IN THE RE FORMAT

L2 ANSWER 11 OF 40 HCAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 2001:850956 HCAPLUS  
DOCUMENT NUMBER: 135:376777  
TITLE: Peptide pharmaceutical formulations  
INVENTOR(S): Holmquist, Barton; Dormady, Daniel C.  
PATENT ASSIGNEE(S): Bionebraska, Inc., USA  
SOURCE: PCT Int. Appl., 35 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001087322	A2	20011122	WO 2001-US15872	20010517
WO 2001087322	A3	20020718		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
US 2002061838	A1	20020523	US 2001-858880	20010517
PRIORITY APPLN. INFO.:			US 2000-205377P	P 20000517
			US 2000-205262P	P 20000519

AB A pharmaceutical compn. for administration to a mammal is disclosed. The compn. includes a therapeutically effective amt. of a peptide, such as a GLP-1 mol., a PTH mol., or a GRF mol. The compn. further includes a buffer including a weak acid having an acid dissociation const. value of greater than about  $1 \times 10^{-5}$ , such as acetic acid. The compn. also includes an excipient for making the compn. generally isotonic, such as D-mannitol.

IT **141758-74-9**, Exendin 4 (*Heloderma suspectum*)  
RL: PEP (Physical, engineering or chemical process); PRP  
(Properties); THU (Therapeutic use); BIOL (Biological study); PROC  
(Process); USES (Uses)  
(peptide pharmaceutical formulations relating to parathormone,  
glucagon-like peptide-1, and growth hormone-releasing factor)

L2 ANSWER 12 OF 40 HCAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 2001:759575 HCAPLUS

09/756690

DOCUMENT NUMBER: 135:298797  
TITLE: Synergistic effect of a sulfonylurea and/or non-sulfonylurea K<sup>+</sup> ATP channel blocker, and a phosphodiesterase 3 type inhibitor for the treatment of non-insulin-dependent diabetes or other conditions  
INVENTOR(S): Fryburg, David Albert; Parker, Janice Catherine  
PATENT ASSIGNEE(S): Pfizer Products Inc., USA  
SOURCE: Eur. Pat. Appl., 19 pp.  
CODEN: EPXXDW  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 1145717	A2	20011017	EP 2001-303020	20010330
EP 1145717	A3	20020814		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
US 2002013268	A1	20020131	US 2001-829874	20010410
CA 2343850	AA	20011013	CA 2001-2343850	20010411
BR 2001001461	A	20011113	BR 2001-1461	20010411
JP 2001354568	A2	20011225	JP 2001-115674	20010413

PRIORITY APPLN. INFO.: US 2000-196728P P 20000413  
AB The invention provides the use of a synergistic amt. of (1) a sulfonylurea, a non-sulfonylurea K<sup>+</sup> ATP channel blocker, or a sulfonylurea and a non-sulfonylurea K<sup>+</sup> ATP channel blocker; and (2) a cAMP phosphodiesterase type 3 inhibitor; for the manuf. of medicaments for treating or preventing non-insulin-dependent diabetes mellitus, insulin resistance, Syndrome X, diabetic neuropathy, diabetic nephropathy, diabetic retinopathy, diabetic cardiomyopathy, polycystic ovary syndrome, cataracts, hyperglycemia, or impaired glucose tolerance. The invention also provides kits and pharmaceutical compns. that comprise (1) a sulfonylurea, a non-sulfonylurea K<sup>+</sup> ATP channel blocker, or a sulfonylurea and a non-sulfonylurea K<sup>+</sup> ATP channel blocker; and (2) a cAMP phosphodiesterase type 3 inhibitor. The invention further provides kits and pharmaceutical compns. that comprise (1) a sulfonylurea, a non-sulfonylurea K<sup>+</sup> ATP channel blocker, or a sulfonylurea and a non-sulfonylurea K<sup>+</sup> ATP channel blocker; (2) a cAMP phosphodiesterase type 3 inhibitor; and (3) an addnl. compd. useful for the treatment of non-insulin-dependent diabetes mellitus, insulin resistance, Syndrome X, diabetic neuropathy, diabetic nephropathy, diabetic retinopathy, diabetic cardiomyopathy, polycystic ovary syndrome, cataracts, hyperglycemia, or impaired glucose tolerance.

IT 335149-21-8, AC2993

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(sulfonylurea and/or non-sulfonylurea K<sup>+</sup> ATP channel blocker and phosphodiesterase 3 type inhibitor synergism for treatment of non-insulin-dependent diabetes or other conditions)

L2 ANSWER 13 OF 40 HCAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 2001:650487 HCAPLUS

Searcher : Shears 308-4994

DOCUMENT NUMBER: 135:205920  
 TITLE: Metabolic intervention with GLP-1 to improve the  
 function of ischemic and reperfused tissue  
 INVENTOR(S): Coolidge, Thomas R.; Ehlers, Mario R. W.  
 PATENT ASSIGNEE(S): BioNebraska, Inc., USA  
 SOURCE: U.S., 10 pp.  
 CODEN: USXXAM  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 3  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 6284725	B1	20010904	US 1999-302596	19990430
WO 2000066138	A2	20001109	WO 2000-US11251	20000427
WO 2000066138	A3	20010705		
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
EP 1173197	A2	20020123	EP 2000-926404	20000427
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
JP 2002543142	T2	20021217	JP 2000-615022	20000427
US 2002055460	A1	20020509	US 2001-851738	20010509
US 2002147131	A1	20021010	US 2001-953021	20010911
NO 2001005294	A	20011228	NO 2001-5294	20011029
PRIORITY APPLN. INFO.: US 1998-103498P P 19981008 US 1999-302596 A 19990430 WO 2000-US11251 W 20000427 US 2001-851738 A1 20010509				

AB Individuals in need of treatment of ischemia-related reperfusion are treated, preferably i.v., with a compn. which includes a compd. which binds to a receptor for the glucagon-like peptide-1. The invention relates to both the method and compns. for such treatment.

IT 203743-40-2

RL: PRP (Properties)

(unclaimed protein sequence; metabolic intervention with GLP-1 to improve the function of ischemic and reperfused tissue)

REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 14 OF 40 HCAPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 2001:525943 HCAPLUS  
 DOCUMENT NUMBER: 135:132445  
 TITLE: Use of exendins and agonists thereof for modulation of triglyceride levels and treatment of dyslipidemia  
 INVENTOR(S): Kolterman, Orville Gene; Young, Andrew A.  
 PATENT ASSIGNEE(S): Amylin Pharmaceuticals, Inc., USA  
 SOURCE: PCT Int. Appl., 161 pp.

09/756690

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

3

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001051078	A1	20010719	WO 2001-US719	20010109
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
EP 1246638	A1	20021009	EP 2001-900978	20010109
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR				
PRIORITY APPLN. INFO.:			US 2000-175365P	P 20000110
			WO 2001-US719	W 20010109

AB Methods for modulating the levels of plasma triglyceride and other lipids in a subject comprise administration of an effective amt. of an exendin or exendin agonist, alone or in conjunction with other compds. or compns. that lower blood triglyceride and/or other lipid levels.

IT 203743-40-2

RL: PRP (Properties)

(unclaimed protein sequence; use of exendins and agonists thereof for modulation of triglyceride levels and treatment of dyslipidemia)

REFERENCE COUNT:

7

THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 15 OF 40 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2001:283949 HCAPLUS

DOCUMENT NUMBER: 134:311218

TITLE: Synthesis and use of heterocyclic sodium/proton exchange inhibitors

INVENTOR(S): Ahmad, Saleem; Wu, Shung C.; O'Neil, Steven V.; Ngu, Khehyong; Atwal, Karnail S.

PATENT ASSIGNEE(S): Bristol-Myers Squibb Company, USA

SOURCE: PCT Int. Appl., 221 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

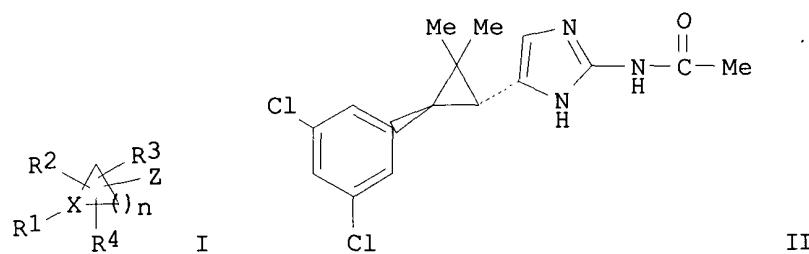
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001027107	A2	20010419	WO 2000-US27461	20001002
WO 2001027107	A3	20020124		

Searcher : Shears 308-4994

09/756690

OTHER SOURCE(S): MARPAT 134:311218  
GI



AB Compds. of formula I [wherein; n is 1-5; X is N or CR<sub>5</sub>, where R<sub>5</sub> is H, halo, alkenyl, alkynyl, alkoxy, alkyl, aryl or heteroaryl; Z is a heteroaryl group; R<sub>1</sub> is H, alk(en)(yn)yl, alk(enyl)(ynyl)oxy, (aryl or alkyl)3Si, cycloalk(en)yl, (aryl)amino, aryl(alkyl), cycloheteroaryl, etc.; R<sub>2</sub>, R<sub>3</sub> and R<sub>4</sub> are any of the groups set out for R<sub>1</sub> and optionally substituted with 1 to 5 substituents which may be the same or different and when X is N, R<sub>1</sub> is preferably aryl or heteroaryl] are claimed. Several hundred examples are disclosed. Synthesis of II proceeds via cyclopropanation of the cinnamate derived from the olefination between 3,5-dichlorobenzaldehyde and t-butyldiethylphosphonoacetate. The intermediate tert-Bu ester is converted to the corresponding .alpha.-chloroketone and reacted with acetyl guanidine to provide II in a total of 5 steps. Compds. I are said to be sodium/proton exchange inhibitors (NHE). Pharmaceutical combinations are claimed using I and certain antihypertensive agents, .beta.-adrenergic agonists, hypolipidemic agents, antidiabetic agents, antiobesity agents, etc. Compds. I are useful as antianginal and cardioprotective agents and provide a method for preventing or treating angina pectoris, cardiac dysfunction, myocardial necrosis, and arrhythmia.

IT 335149-21-8, AC 2993

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

09/756690

(pharmaceuticals also contg.; synthesis and use of heterocyclic sodium/proton exchange inhibitors)

L2 ANSWER 16 OF 40 HCPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 2001:247368 HCPLUS  
DOCUMENT NUMBER: 134:290749  
TITLE: Pituitary adenylate cyclase activating peptide (PACAP) receptor 3 (R3) agonists and their pharmacological methods of use in treating metabolic disorders and respiratory disease  
INVENTOR(S): Pan, Clark; Tsutsumi, Manami; Shanafelt, Armen B.  
PATENT ASSIGNEE(S): Bayer Corporation, USA  
SOURCE: PCT Int. Appl., 62 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001023420	A2	20010405	WO 2000-US26638	20000927
WO 2001023420	A3	20010830		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
EP 1192182	A2	20020403	EP 2000-967002	20000927
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
PRIORITY APPLN. INFO.:			US 1999-407832	A 19990928
			US 2000-595280	A 20000615
			WO 2000-US26638	W 20000927

AB The invention provides novel peptides that function *in vivo* to stimulate insulin release from pancreatic beta cells in a glucose-dependent fashion. These insulin secretagogue peptides are shown to stimulate insulin release in rat islet cells *in vitro*, and *in vivo*. The peptides of the present invention provide a new therapy for patients with decreased endogenous insulin secretion, in particular type 2 diabetics. In particular, the invention is a polypeptide selected from a specific group of VIP/PACAP-related polypeptides, or functional equiv. thereof. The invention is also directed to a method of treating a metabolic disease or a respiratory disease in a mammal comprising administering a therapeutically effective amt. of the insulin secretagogue peptides to said mammal. Also disclosed are methods of making the peptides, both recombinant and synthetic; pharmaceutical compns. contg. the peptides; and antibodies to the peptides.

IT 203743-40-2

RL: PRP (Properties)

(unclaimed protein sequence; pituitary adenylate cyclase

09/756690

activating peptide (PACAP) receptor 3 (R3) agonists and their pharmacol. methods of use in treating metabolic disorders and respiratory disease)

L2 ANSWER 17 OF 40 HCAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 2001:50681 HCAPLUS  
DOCUMENT NUMBER: 134:110470  
TITLE: Preparation of stable peptide conjugates containing variants of exendin-4 and GLP-I that lower blood glucose levels and regulate gastric emptying  
INVENTOR(S): Larsen, Bjarne Due; Mikkelsen, Jens Damsgaard; Neve, Soren  
PATENT ASSIGNEE(S): Zealand Pharmaceuticals A/S, Den.  
SOURCE: PCT Int. Appl., 83 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001004156	A1	20010118	WO 2000-DK393	20000712
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
EP 1076066	A1	20010214	EP 1999-610043	19990809
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
EP 1196444	A1	20020417	EP 2000-945656	20000712
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
JP 2003505347	T2	20030212	JP 2001-509765	20000712
PRIORITY APPLN. INFO.:			US 1999-143591P P	19990712
			EP 1999-610043 A	19990809
			WO 2000-DK393 W	20000712

OTHER SOURCE(S): MARPAT 134:110470  
AB The present invention relates to novel X-Z peptide conjugates which have increased stability and are useful in the treatment of excess levels of blood glucose. Peptide X selected from the group consisting of (a) an exendin having at least 90% homol. to exendin-4; (b) a variant of said exendin wherein said variant comprises a modification selected from the group consisting of between one and five deletions at positions 34-39 and contains a Lys at position 40 having a lipophilic substituent; or (c) GLP-I (7-36) or GLP-I (7-37) having at least one modification selected from the group consisting of: (i) substitution of D-alanine, glycine or alpha-amino isobutyric acid for alanine at position 8 and (ii) a lipophilic substituent. Peptide Z is a peptide sequence of 4-20 amino acid units covalently bound to variant X, wherein each amino acid unit in said peptide sequence, Z, is selected from the group

consisting of Ala, Leu, Ser, Thr, Tyr, Asn, Gln, Asp, Glu, Lys, Arg, His, Met, Orn, and amino acid units of the general formula -NH-C(R1)(R2)-C(=O)- wherein R1 and R2 are selected from the group consisting of hydrogen, C1-6-alkyl, Ph, and phenylmethyl, which are optionally substituted. R1 and R2 together with the carbon atom to which they are bound can also form a cyclopentyl, cyclohexyl, or cycloheptyl ring. The peptide X is further characterized in being effective in improving glucose tolerance in a diabetic mammal. The peptides are effective in the treatment of diseases that benefit from regulation of excess levels of blood glucose and /or regulation of gastric emptying, such as diabetes and eating disorders. The present invention also relates to methods of prep. said novel peptides and pharmaceutical compns. contg. the peptides.

IT

**320367-11-1P 320367-31-5P**

RL: BAC (Biological activity or effector, except adverse); BPN (Biosynthetic preparation); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of stable peptide conjugates contg. variants of exendin-4 and GLP-I that lower blood glucose levels and regulate gastric emptying)

REFERENCE COUNT:

11 THERE ARE 11 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 18 OF 40 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2000:900675 HCAPLUS

DOCUMENT NUMBER: 134:51920

TITLE: GLP-1 as a diagnostic test to determine .beta.-cell function and the presence of impaired glucose tolerance (IGT) and type-II diabetes

INVENTOR(S): Holst, J. J.; Vilsboll, Tina

PATENT ASSIGNEE(S): Bioneerbraska, Inc., USA

SOURCE: PCT Int. Appl., 42 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000077039	A2	20001221	WO 2000-US16428	20000614
WO 2000077039	A3	20010329		
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
US 6344180	B1	20020205	US 1999-333415	19990615
EP 1185308	A2	20020313	EP 2000-939881	20000614
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				

PRIORITY APPLN. INFO.: US 1999-333415 A 19990615  
WO 2000-US16428 W 20000614

AB Since glucagon-like peptide-1 (GLP-1) is the most potent insulinotropic hormone known and has been shown to stimulate insulin secretion strongly in patients with type II diabetes, this invention uses GLP-1 or its biol. active analogs in .beta.-cell stimulatory tests in order to test .beta.-cell function in a simple way. The test provides information about insulin secretory capacity, is easy and reproducible and has insignificant side effects.

IT **203743-40-2**

RL: PRP (Properties)  
(unclaimed protein sequence; gLP-1 as a diagnostic test to det. .beta.-cell function and the presence of impaired glucose tolerance (IGT) and type-II diabetes)

L2 ANSWER 19 OF 40 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2000:861704 HCAPLUS

DOCUMENT NUMBER: 134:37033

TITLE: Use of exendins and agonists thereof for the treatment of gestational diabetes mellitus

INVENTOR(S): Hiles, Richard; Prickett, Kathryn S.

PATENT ASSIGNEE(S): Amylin Pharmaceuticals, Inc., USA

SOURCE: PCT Int. Appl., 133 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000073331	A2	20001207	WO 2000-US14231	20000523
WO 2000073331	A3	20010628		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
US 6506724	B1	20030114	US 1999-323867	19990601
EP 1181043	A2	20020227	EP 2000-937710	20000523
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
JP 2003501361	T2	20030114	JP 2001-500655	20000523

PRIORITY APPLN. INFO.: US 1999-323867 A 19990601  
WO 2000-US14231 W 20000523

AB Methods for treating gestational diabetes which comprise administration of an effective amt. of an exendin or an exendin agonist, alone or in conjunction with other compds. or compns. that lower blood glucose levels.

IT **210829-56-4P 210829-59-7P**

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

09/756690

(use of exendins and agonists thereof for treatment of  
gestational diabetes mellitus in relation to combination with  
insulin or amylin agonist)

IT 141758-74-9, Exendin 4 (Heloderma suspectum)

203743-40-2

RL: BAC (Biological activity or effector, except adverse); BSU  
(Biological study, unclassified); THU (Therapeutic use); BIOL  
(Biological study); USES (Uses)

(use of exendins and agonists thereof for treatment of  
gestational diabetes mellitus in relation to combination with  
insulin or amylin agonist)

L2 ANSWER 20 OF 40 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2000:824301 HCAPLUS

DOCUMENT NUMBER: 134:13338

TITLE: Long lasting insulinotropic peptides

INVENTOR(S): Bridon, Dominique P.; L'Archeveque, Benoit;  
Ezrin, Alan M.; Holmes, Darren L.; Leblanc,  
Anouk; St. Pierre, Serge

PATENT ASSIGNEE(S): Conjuchem, Inc., Can.

SOURCE: PCT Int. Appl., 96 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000069911	A1	20001123	WO 2000-US13563	20000517
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
WO 2000070665	A2	20001123	WO 2000-IB763	20000517
WO 2000070665	A3	20010419		
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW				
RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
EP 1171582	A2	20020116	EP 2000-929748	20000517
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
EP 1180121	A1	20020220	EP 2000-930796	20000517
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
BR 2000010750	A	20020226	BR 2000-10750	20000517

09/756690

AU 754770	B2	20021121	AU 2000-48555	20000517
EP 1264840	A1	20021211	EP 2002-14617	20000517
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL				
JP 2003500341	T2	20030107	JP 2000-619018	20000517
US 6329336	B1	20011211	US 2000-623618	20000905
US 6514500	B1	20030204	US 2000-657332	20000907
US 2002049153	A1	20020425	US 2001-876388	20010606
NO 2001005584	A	20020103	NO 2001-5584	20011115
PRIORITY APPLN. INFO.:				
			US 1999-134406P	P 19990517
			US 1999-159783P	P 19991015
			US 1999-153406P	P 19990910
			EP 2000-932570	A3 20000517
			WO 2000-IB763	W 20000517
			WO 2000-US13563	W 20000517
			US 2000-623618	A3 20000905

AB Modified insulinotropic peptides are disclosed. The modified insulinotropic peptides are capable of forming a peptidase stabilized insulinotropic peptide. The modified insulinotropic peptides are capable of forming covalent bonds with one or more blood components to form a conjugate. The conjugates may be formed in vivo or ex vivo. The modified peptides are administered to treat humans with diabetes and other related diseases.

IT **309729-73-5**

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(lasting insulinotropic pepwith antidiabetic activity)

IT **203743-40-2 308815-99-8**

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(long lasting insulinotropic peptides with antidiabetic activity)

IT **308244-92-0P 309728-25-4P 309729-78-0P**

RL: SPN (Synthetic preparation); PREP (Preparation)  
(long lasting insulinotropic peptides with antidiabetic activity)

REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR  
THIS RECORD. ALL CITATIONS AVAILABLE IN  
THE RE FORMAT

L2 ANSWER 21 OF 40 HCAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 2000:824291 HCAPLUS  
DOCUMENT NUMBER: 134:21425  
TITLE: Protection of endogenous therapeutic peptides  
from peptidase activity through conjugation to  
blood components  
INVENTOR(S): Bridon, Dominique P.; Ezrin, Alan M.; Milner,  
Peter G.; Holmes, Darren L.; Thibaudeau, Karen  
Conjuchem, Inc., Can.  
PATENT ASSIGNEE(S):  
SOURCE: PCT Int. Appl., 733 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 3  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
-----	-----	-----	-----	-----

Searcher : Shears 308-4994

09/756690

WO 2000069900 A2 20001123 WO 2000-US13576 20000517  
WO 2000069900 A3 20010215  
WO 2000069900 C2 20020704

W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR,  
CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,  
ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,  
LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU,  
SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ,  
VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM  
RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,  
DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF,  
BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

WO 2000070665 A2 20001123 WO 2000-IB763 20000517  
WO 2000070665 A3 20010419

W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR,  
CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,  
ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,  
LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU,  
SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ,  
VN, YU, ZA, ZW  
RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AM, AZ, BY, KG,  
KZ, MD, RU, TJ, TM, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB,  
GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA,  
GN, GW, ML, MR, NE, SN, TD, TG

EP 1105409 A2 20010613 EP 2000-936023 20000517  
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC,  
PT, IE, SI, LT, LV, FI, RO  
EP 1171582 A2 20020116 EP 2000-929748 20000517  
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC,  
PT, IE, SI, LT, LV, FI, RO  
EP 1264840 A1 20021211 EP 2002-14617 20000517  
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC,  
PT, IE, SI, LT, LV, FI, RO, MK, CY, AL

JP 2003500341 T2 20030107 JP 2000-619018 20000517  
US 6514500 B1 20030204 US 2000-657332 20000907

PRIORITY APPLN. INFO.: US 1999-134406P P 19990517  
US 1999-153406P P 19990910  
US 1999-159783P P 19991015  
EP 2000-932570 A3 20000517  
WO 2000-IB763 W 20000517  
WO 2000-US13576 W 20000517

AB A method for protecting a peptide from peptidase activity in vivo, the peptide being composed of between 2 and 50 amino acids and having a C-terminus and an N-terminus and a C-terminus amino acid and an N-terminus amino acid is described. In the first step of the method, the peptide is modified by attaching a reactive group to the C-terminus amino acid, to the N-terminus amino acid, or to an amino acid located between the N-terminus and the C-terminus, such that the modified peptide is capable of forming a covalent bond in vivo with a reactive functionality on a blood component. The solid phase peptide synthesis of a no. of derivs. with 3-maleimidopropionic acid (3-MPA) is described. In the next step, a covalent bond is formed between the reactive group and a reactive functionality on a blood component to form a peptide-blood component conjugate, thereby protecting said peptide from peptidase activity. The final step of the method involves the analyzing of the stability of the peptide-blood component conjugate to assess the protection of the peptide from peptidase activity. Thus, the percentage of a K5

kringle peptide (Pro-Arg-Lys-Leu-Tyr-Asp-Lys-NH<sub>2</sub>) conjugated to human serum albumin via MPA remained relatively const. through a 24-h plasma assay in contrast to unmodified K5 which decreased to 9% of the original amt. of K5 in only 4 h in plasma.

IT 308245-14-9P 308245-46-7P

RL: RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(protection of endogenous therapeutic peptides from peptidase activity through conjugation to blood components)

IT 203743-40-2 309257-15-6

RL: PRP (Properties)

(unclaimed protein sequence; protection of endogenous therapeutic peptides from peptidase activity through conjugation to blood components)

L2 ANSWER 22 OF 40 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2000:790546 HCAPLUS

DOCUMENT NUMBER: 133:359242

TITLE: Modified exendins and exendin agonists

INVENTOR(S): Young, Andrew; Prickett, Kathryn

PATENT ASSIGNEE(S): Amylin Pharmaceuticals, Inc., USA

SOURCE: PCT Int. Appl., 119 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000066629	A1	20001109	WO 2000-US11814	20000428
W:	AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
EP 1175443	A1	20020130	EP 2000-928685	20000428
R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO			
BR 2000010705	A	20020205	BR 2000-10705	20000428
JP 2002544127	T2	20021224	JP 2000-615657	20000428
PRIORITY APPLN. INFO.:			US 1999-132018P	P 19990430
			WO 2000-US11814	W 20000428

AB Novel modified exendins and exendin agonists having an exendin or exendin agonist linked to one or more polyethylene glycol polymers, for example, and related formulations and dosages and methods of administration thereof are provided. These modified exendins and exendin agonists, compns. and methods are useful in treating diabetes and conditions that would be benefited by lowering plasma glucose or delaying and/or slowing gastric emptying or inhibiting food intake.

IT 305814-59-9P 305815-28-5P 305815-30-9P  
305818-90-0P

09/756690

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PNU (Preparation, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
(modified exendins and exendin agonists)

IT **141758-74-9**, Exendin 4 (*Heloderma suspectum*)

RL: PRP (Properties)

(unclaimed sequence; modified exendins and exendin agonists)

REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 23 OF 40 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2000:790326 HCAPLUS

DOCUMENT NUMBER: 133:345167

TITLE: Metabolic intervention with GLP-1 or its biologically active analogues to improve the function of the ischemic and reperfused brain

INVENTOR(S): Coolidge, Thomas R.; Ehlers, Mario R. W.

PATENT ASSIGNEE(S): Bioneerbraska, Inc., USA

SOURCE: PCT Int. Appl., 19 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000066142	A2	20001109	WO 2000-US11652	20000501
WO 2000066142	A3	20020124		
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
US 6429197	B1	20020806	US 1999-303016	19990430
EP 1187628	A2	20020320	EP 2000-928616	20000501
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
JP 2002543145	T2	20021217	JP 2000-615026	20000501
NO 2001005298	A	20011228	NO 2001-5298	20011029
PRIORITY APPLN. INFO.:			US 1999-303016	A 19990430
			US 1998-103498P	P 19981008
			WO 2000-US11652	W 20000501

AB It has now been discovered that GLP-1 treatment after acute stroke or hemorrhage, preferably i.v. administration, can be an ideal treatment because it provides a means for optimizing insulin secretion, increasing brain anabolism, enhancing insulin effectiveness by suppressing glucagon, and maintaining euglycemia or mild hypoglycemia with no risk of severe hypoglycemia.

IT **203743-40-2**

RL: PRP (Properties)

(unclaimed protein sequence; metabolic intervention with GLP-1 or

09/756690

its biol. active analogs to improve the function of the ischemic and reperfused brain)

L2 ANSWER 24 OF 40 HCAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 2000:790323 HCAPLUS  
DOCUMENT NUMBER: 133:345166  
TITLE: Metabolic intervention with GLP-1 to improve the function of ischemic and reperfused tissue  
INVENTOR(S): Coolidge, Thomas R.; Ehlers, Mario R. W.  
PATENT ASSIGNEE(S): Bioneerbraska, Inc., USA  
SOURCE: PCT Int. Appl., 22 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 3  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000066138	A2	20001109	WO 2000-US11251	20000427
WO 2000066138	A3	20010705		
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
US 6284725	B1	20010904	US 1999-302596	19990430
EP 1173197	A2	20020123	EP 2000-926404	20000427
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
JP 2002543142	T2	20021217	JP 2000-615022	20000427
NO 2001005294	A	20011228	NO 2001-5294	20011029
PRIORITY APPLN. INFO.:			US 1999-302596 A	19990430
			US 1998-103498P P	19981008
			WO 2000-US11251 W	20000427

AB Individuals in need of treatment of ischemia-related reperfusion are treated, preferably i.v., with a compn. which includes a compd. which binds to a receptor for the glucagon-like peptide-1. The invention relates to both the method and compns. for such treatment.

IT 203743-40-2

RL: PRP (Properties)  
(unclaimed protein sequence; metabolic intervention with GLP-1 to improve the function of ischemic and reperfused tissue)

L2 ANSWER 25 OF 40 HCAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 2000:493318 HCAPLUS  
DOCUMENT NUMBER: 133:129880  
TITLE: Methods using an exendin or related substance for glucagon suppression  
INVENTOR(S): Young, Andrew; Gedulin, Bronislava  
PATENT ASSIGNEE(S): Amylin Pharmaceuticals, Inc., USA  
SOURCE: PCT Int. Appl., 96 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

## PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000041548	A2	20000720	WO 2000-US942	20000114
WO 2000041548	A3	20001130		
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
CA 2356331	AA	20000720	CA 2000-2356331	20000114
EP 1143989	A2	20011017	EP 2000-902415	20000114
EP 1143989	A3	20020911		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
BR 2000007823	A	20011120	BR 2000-7823	20000114
JP 2002538084	T2	20021112	JP 2000-593169	20000114
NO 2001003469	A	20010914	NO 2001-3469	20010712
PRIORITY APPLN. INFO.:			US 1999-116380P	P 19990114
			US 1999-132017P	P 19990430
			US 2000-175365P	P 20000110
			WO 2000-US942	W 20000114

AB Methods are provided for use of an exendin, an exendin agonist, or a modified exendin or exendin agonist having an exendin or exendin agonist linked to one or more polyethylene glycol polymers, for example, for lowering glucagon levels and/or suppressing glucagon secretion in a subject. These methods are useful in treating hyperglucagonemia and other conditions that would be benefited by lowering plasma glucagon or suppressing glucagon secretion.

IT **141758-74-9P**, Exendin 4 (*Heloderma suspectum*)  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
 (exendin or related substance for glucagon suppression)

IT **141758-74-9**, Exendin 4 (*Heloderma suspectum*)  
 RL: PRP (Properties)  
 (unclaimed protein sequence; methods using an exendin or related substance for glucagon suppression)

L2 ANSWER 26 OF 40 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2000:493315 HCAPLUS

DOCUMENT NUMBER: 133:135612

TITLE: Novel exendin agonist formulations and methods of administration thereof

INVENTOR(S): Young, Andrew; L'Italien, James J.; Kolterman, Orville

PATENT ASSIGNEE(S): Amylin Pharmaceuticals, Inc., USA

SOURCE: PCT Int. Appl., 281 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

09/756690

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000041546	A2	20000720	WO 2000-US902	20000114
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
CA 2356706	AA	20000720	CA 2000-2356706	20000114
EP 1140145	A2	20011010	EP 2000-914425	20000114
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
BR 2000007820	A	20011120	BR 2000-7820	20000114
JP 2002534450	T2	20021015	JP 2000-593167	20000114
NO 2001003468	A	20010914	NO 2001-3468	20010712
PRIORITY APPLN. INFO.:			US 1999-116380P	P 19990114
			US 2000-175365P	P 20000110
			WO 2000-US902	W 20000114

AB Novel exendin and exendin agonist compd. formulations and dosages and methods of administration thereof are provided. These compns. and methods are useful in treating diabetes and conditions that would be benefited by lowering plasma glucose or delaying and/or slowing gastric emptying or inhibiting food intake.

IT **141758-74-9P**, Exendin-4 (Heloderma suspectum)

RL: BAC (Biological activity or effector, except adverse); BOC (Biological occurrence); BSU (Biological study, unclassified); PNU (Preparation, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); OCCU (Occurrence); PREP (Preparation); USES (Uses)  
(amino acid sequence; novel exendin agonist formulations and methods of administration thereof as antidiabetic agents and appetite suppressants)

L2 ANSWER 27 OF 40 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2000:133809 HCAPLUS

DOCUMENT NUMBER: 132:175839

TITLE: Differentiation of non-insulin producing cells into insulin producing cells by GLP-1 or Exendin-4 and uses thereof

INVENTOR(S): Egan, Josephine; Perfetti, Riccardo; Passaniti, Antonino; Greig, Nigel; Holloway, Harold

PATENT ASSIGNEE(S): United States of America, Department of Health and Human Services, USA

SOURCE: PCT Int. Appl., 119 pp.  
CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

Searcher : Shears 308-4994

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000009666	A2	20000224	WO 1999-US18099	19990810
WO 2000009666	A3	20001123		
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
CA 2339326	AA	20000224	CA 1999-2339326	19990810
AU 9955524	A1	20000306	AU 1999-55524	19990810
EP 1105460	A2	20010613	EP 1999-942066	19990810
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI				
PRIORITY APPLN. INFO.:			US 1998-95917P	P 19980810
			WO 1999-US18099	W 19990810

AB The present invention relates to a population of insulin producing cells made by a process comprising contacting non-insulin producing cells with a growth factor selected from the group consisting of GLP-1 or Exendin-4, growth factors having amino acid sequences substantially homologous to GLP-1 or Exendin-4, and fragments thereof. The present invention also relates to methods of differentiating non-insulin producing cells into insulin producing cells and of enriching a population of cells for insulin-producing cells. The present invention also relates to methods of treating diabetes. Exendin-4 was more potent an insulinotropic agent than GLP-1 on several levels when given i.v.

IT **203743-40-2**

RL: PRP (Properties)  
(unclaimed protein sequence; differentiation of non-insulin producing cells into insulin producing cells by GLP-1 or Exendin-4 and uses thereof)

L2 ANSWER 28 OF 40 HCPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 1999:18104 HCPLUS  
DOCUMENT NUMBER: 130:178590  
TITLE: Black widow spider .alpha.-latrotoxin: a presynaptic neurotoxin that shares structural homology with the glucagon-like peptide-1 family of insulin secretagogic hormones  
AUTHOR(S): Holz, George G.; Habener, Joel F.  
CORPORATE SOURCE: Diabetes Unit, Howard Hughes Medical Institute, Massachusetts General Hospital, Harvard Medical School, Boston, MA, 02114; USA  
SOURCE: Comparative Biochemistry and Physiology, Part B: Biochemistry & Molecular Biology (1998), 121B(2), 177-184  
PUBLISHER: Elsevier Science Inc.  
DOCUMENT TYPE: Journal  
LANGUAGE: English  
AB .alpha.-Latrotoxin is a presynaptic neurotoxin isolated from the venom of the black widow spider *Latrodectus tredecimguttatus*. It exerts toxic effects in the vertebrate central nervous system by

09/756690

depolarizing neurons, by increasing  $[Ca^{2+}]_i$  and by stimulating uncontrolled exocytosis of neurotransmitters from nerve terminals. The actions of .alpha.-latrotoxin are mediated, in part, by a GTP-binding protein-coupled receptor referred to as CIRL or latrophilin. Exendin-4 is also a venom toxin, and it is derived from the salivary gland of the Gila monster *Heloderma suspectum*. It acts as an agonist at the receptor for glucagon-like peptide-1(7-36)-amide (GLP-1), thereby stimulating secretion of insulin from pancreatic .beta.-cells of the islets of Langerhans. Here is reported a surprising structural homol. between a-latrotoxin and exendin-4 that is also apparent amongst all members of the GLP-1-like family of secretagogic hormones (GLP-1, glucagon, vasoactive intestinal polypeptide, secretin, pituitary adenylyl cyclase activating polypeptide). On the basis of this homol., we report the synthesis and initial characterization of a chimeric peptide (Black Widow GLP-1) that stimulates  $Ca^{2+}$  signaling and insulin secretion in human .beta.-cells and MIN6 insulinoma cells. It is also reported here that the GTP-binding protein-coupled receptors for .alpha.-latrotoxin and exendin-4 share highly significant structural similarity in their extracellularly-oriented amino-termini. We propose that mol. mimicry has generated conserved structural motifs in secretagogic toxins and their receptors, thereby explaining the evolution of defense or predatory strategies that are shared in common amongst distantly related species including spiders, lizards, and snakes. Evidently, the toxic effects of .alpha.-latrotoxin and exendin-4 are explained by their ability to interact with GTP-binding protein-coupled receptors that normally mediate the actions of endogenous hormones or neuropeptides.

IT 141758-74-9, Exendin 4 (*Heloderma suspectum*)

RL: PRP (Properties)

(latrotoxin shares structural homol. with glucagon-like peptide-1 family of insulin secretagogic hormones)

REFERENCE COUNT: 40 THERE ARE 40 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 29 OF 40 HCAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 1998:550504 HCAPLUS  
DOCUMENT NUMBER: 129:185369  
TITLE: Polynucleotides encoding proexendin, and methods and uses thereof  
INVENTOR(S): Drucker, Daniel J.  
PATENT ASSIGNEE(S): 1149336 Ontario Inc., Can.  
SOURCE: PCT Int. Appl., 27 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9835033	A1	19980813	WO 1998-CA71	19980204
W:	AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL,			

Searcher : Shears 308-4994

09/756690

TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ,  
MD, RU, TJ, TM  
RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES,  
FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG,  
CI, CM, GA, GN, ML, MR, NE, SN, TD, TG  
AU 9858507 A1 19980826 AU 1998-58507 19980204  
EP 981611 A1 20000301 EP 1998-901908 19980204  
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC,  
PT, IE, FI  
JP 2001512307 T2 20010821 JP 1998-533455 19980204  
PRIORITY APPLN. INFO.: US 1997-37412P P 19970205  
GB 1997-2582 A 19970207  
WO 1998-CA71 W 19980204

AB Exendin 4 is a biol. active peptide first isolated from Gila monster venom. The invention encompasses polynucleotides encoding proexendin peptides, including exendin and novel peptides, as well as isolated or recombinant proexendin peptides. The invention also includes antibodies which specifically recognize such peptides.  
IT 211430-73-8, Exendin ENTP (Heloderma horridum)  
RL: BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
(amino acid sequence of mature; gene encoding proexendin from Heloderma horridum and applications)  
IT 188265-76-1, Exendin 4, pro- (Heloderma suspectum)  
203743-40-2 211430-62-5  
RL: BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
(amino acid sequence; gene encoding proexendin from Heloderma horridum and applications)  
REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 30 OF 40 HCAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 1998:490528 HCAPLUS  
DOCUMENT NUMBER: 129:149256  
TITLE: Preparation of exendin peptides for the reduction of food intake  
INVENTOR(S): Beeley, Nigel Robert Arnold; Prickett, Kathryn S.; Bhavsar, Sunil  
PATENT ASSIGNEE(S): Amylin Pharmaceuticals, Inc., USA  
SOURCE: PCT Int. Appl., 214 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9830231	A1	19980716	WO 1998-US449	19980107
W:	AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES,			

Searcher : Shears 308-4994

09/756690

FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG,  
CI, CM, GA, GN, ML, MR, NE, SN, TD, TG

AU 9862394 A1 19980803 AU 1998-62394 19980107  
AU 739020 B2 20011004  
EP 996459 A1 20000503 EP 1998-904545 19980107  
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC,  
PT, IE, FI  
JP 2002508742 T2 20020319 JP 1998-531147 19980107  
US 2002137666 A1 20020926 US 1998-3869 19980107  
WO 9907404 A1 19990218 WO 1998-US16387 19980806  
W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ,  
DE, DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP,  
KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK,  
MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL,  
TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG,  
KZ, MD, RU, TJ, TM  
RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK,  
ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,  
CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG  
AU 9887729 A1 19990301 AU 1998-87729 19980806  
AU 749914 B2 20020704  
EP 1019077 A1 20000719 EP 1998-939260 19980806  
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC,  
PT, IE, FI  
BR 9811866 A 20000815 BR 1998-11866 19980806  
JP 2001513512 T2 20010904 JP 2000-506993 19980806  
CA 2309356 AA 19990527 CA 1998-2309356 19981113  
CA 2310097 AA 19990527 CA 1998-2310097 19981113  
WO 9925727 A2 19990527 WO 1998-US24210 19981113  
W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ,  
DE, DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP,  
KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK,  
MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL,  
TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG,  
KZ, MD, RU, TJ, TM  
RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK,  
ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,  
CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG  
WO 9925728 A1 19990527 WO 1998-US24273 19981113  
W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ,  
DE, DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP,  
KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK,  
MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL,  
TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG,  
KZ, MD, RU, TJ, TM  
RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK,  
ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,  
CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG  
AU 9914046 A1 19990607 AU 1999-14046 19981113  
AU 9914588 A1 19990607 AU 1999-14588 19981113  
EP 1032587 A1 20000906 EP 1998-958573 19981113  
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC,  
PT, IE, FI  
BR 9814189 A 20001003 BR 1998-14189 19981113  
BR 9815670 A 20001017 BR 1998-15670 19981113  
EP 1066314 A1 20010110 EP 1998-957897 19981113  
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC,  
PT, IE, FI

09/756690

JP 2001523688  
PRIORITY APPLN. INFO.:

T2 20011127

JP 2000-521108 19981113  
US 1997-34905P P 19970107  
US 1997-55404P P 19970808  
US 1997-65442P P 19971114  
US 1997-66029P P 19971114  
WO 1998-US449 W 19980107  
WO 1998-US16387 W 19980806  
WO 1998-US24210 W 19981113  
WO 1998-US24273 W 19981113

AB Methods for treating conditions or disorders which can be alleviated by reducing food intake are disclosed which comprise administration of an effective amt. of an exendin or an exendin agonist, alone or in conjunction with other compds. or compns. that effect satiety. Approx. 180 exendin-related peptides were synthesized by the solid-phase method.

IT 203743-40-2P 210829-56-4P 210829-59-7P

RL: FFD (Food or feed use); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); USES (Uses)  
(prepn. of exendin peptides for the redn. of food intake)

REFERENCE COUNT: 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 31 OF 40 HCAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 1998:287874 HCAPLUS  
DOCUMENT NUMBER: 129:78077  
TITLE: Molecular cloning of the helodermin and exendin-4 cDNAs in the lizard. Relationship to vasoactive intestinal polypeptide/pituitary adenylate cyclase activating polypeptide and glucagon-like peptide 1 and evidence against the existence of mammalian homologues  
AUTHOR(S): Pohl, Markus; Wank, Stephen A.  
CORPORATE SOURCE: Digestive Diseases Branch, NIDDK, Natl. Inst. of Health, Bethesda, MD, 20892, USA  
SOURCE: Journal of Biological Chemistry (1998), 273(16), 9778-9784  
CODEN: JBCHA3; ISSN: 0021-9258  
PUBLISHER: American Society for Biochemistry and Molecular Biology  
DOCUMENT TYPE: Journal  
LANGUAGE: English

AB Helodermin and exendin-4, two peptides isolated from the salivary gland of the Gila monster, *Heloderma suspectum*, are approx. 50% homologous to vasoactive intestinal peptide (VIP) and glucagon-like peptide-1 (GLP-1), resp., and interact with the mammalian receptors for VIP and GLP-1 with equal or higher affinity and efficacy. Immunohistochem. studies suggested the presence of helodermin-like peptides in mammals. To det. whether helodermin and exendin-4 are present in mammals and their evolutionary relationship to VIP and GLP-1, their cDNAs were first cloned from Gila monster salivary gland. Northern blots and reverse transcription-polymerase chain reaction of multiple Gila monster tissues identified .apprx.500-base pair transcripts only from salivary gland. Both helodermin and exendin-4 full-length cDNAs were .apprx.500 base pairs long, and they encoded precursor proteins contg. the entire amino acid sequence of helodermin and exendin-4, as well as a 44- or 45-amino acid N-terminal extension peptide, resp., having .apprx.60% homol.

The size and structural organization of these cDNAs indicated that they are closely related to one another but markedly different from known cDNAs for the VIP/GLP-1 peptide family previously identified in both lower and higher evolved species. Cloning of the Gila monster VIP/peptide histidine isoleucine, pituitary adenylate cyclase activating polypeptide, and glucagon/GLP-1 cDNAs and Southern blotting of Gila monster DNA demonstrate the coexistence of sep. genes for these peptides and suggests, along with the restricted salivary gland expression, that helodermin and exendin-4 coevolved to serve a sep. specialized function. Probing of a variety of rat and human tissues on Northern blots, human and rat Southern blots, and genomic and cDNA libraries with either helodermin- or exendin-4-specific cDNAs failed to identify evidence for mammalian homologs. These data indicate that helodermin and exendin-4 are not the precursors to VIP and GLP-1 and that they belong to a sep. peptide family encoded by sep. genes. Furthermore, the existence of as yet undiscovered mammalian homologs to helodermin and exendin-4 seems unlikely.

IT 141758-74-9, Exendin 4 (Heloderma suspectum)  
 188265-76-1, Exendin 4, pro- (Heloderma suspectum)  
 RL: BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
 (amino acid sequence; mol. cloning and sequence of the helodermin and exendin-4 cDNAs in the Gila monster)

REFERENCE COUNT: 45 THERE ARE 45 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 32 OF 40 HCAPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 1998:112250 HCAPLUS  
 DOCUMENT NUMBER: 128:192936  
 TITLE: Preparation of exendin peptide analogs as agonists for regulating gastrointestinal motility  
 INVENTOR(S): Young, Andrew A.; Gedulin, Bronislava; Beeley, Nigel Robert Arnold; Prickett, Kathryn S.  
 PATENT ASSIGNEE(S): Amylin Pharmaceuticals, Inc., USA; Young, Andrew A.; Gedulin, Bronislava; Beeley, Nigel Robert Arnold; Prickett, Kathryn S.  
 SOURCE: PCT Int. Appl., 70 pp.  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9805351	A1	19980212	WO 1997-US14199	19970808
W:	AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, HU, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG			

AU 9740636	A1 19980225	AU 1997-40636	19970808
EP 966297	A1 19991229	EP 1997-938261	19970808
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI			
JP 2001501593	T2 20010206	JP 1998-508263	19970808
PRIORITY APPLN. INFO.:		US 1996-694954 A	19960808
		WO 1997-US14199 W	19970808

OTHER SOURCE(S): MARPAT 128:192936

AB Methods for reducing gastric motility and delaying gastric emptying for therapeutic and diagnostic purposes are disclosed which comprise administration of an effective amt. of an exendin or an exendin agonist H-Xaa1-Xaa2-Xaa3-Gly-Thr-Xaa4-Xaa5-Xaa6-Xaa7-Xaa8-Ser-Lys-Gln-Xaa9-Glu-Glu-Glu-Ala-Val-Arg-Leu-Xaa10-Xaa11-Xaa12-Xaa13-Leu-Lys-Asn-Gly-Gly-Xaa14-Ser-Ser-Gly-Ala-Xaa15-Xaa16-Xaa17-Xaa18-Z [Xaa1 = His, Arg, Tyr; Xaa2 = Ser, Gly, Ala, Thr; Xaa3, Xaa7, Xaa12 = independently Asp, Glu; Xaa4, Xaa10 = independently Phe, Tyr, naphthylalanine; Xaa5, Xaa6 = independently Thr, Ser; Xaa8, Xaa9 = independently Leu, Ile, Val, pentylglycine, Met; Xaa11 = any group Xaa8, tert-butylglycine; Xaa13 = any group Xaa4, Trp; Xaa14-Xaa17 = independently Pro, homoproline, 3-Hyp, 4-Hyp, thioproline, N-alkylglycine, N-alkylpentylglycine, N-alkylalanine; Xaa18 = Ser, Thr, Tyr; Z = OH, NH2; with the proviso that the compd. does not have the formula of exendin-3 or exendin-4] or a pharmaceutically acceptable salt thereof. Methods for treating conditions assocd. with elevated, inappropriate, or undesired post-prandial blood glucose levels are disclosed which comprise administration of an effective amt. of an exendin or an exendin agonist alone or in conjunction with other anti-gastric emptying agents. Thus, exendin-4 acid and [Leu14, Phe25]-exendin-4, prep'd. by std. solid-phase methods on a 4-(2,4-dimethoxyphenyl)-Fmoc-aminomethylphenoxyacetamide norleucine-MBHA resin using 9-fluorenylmethoxycarbonyl (Fmoc)-protected amino acids, inhibited gastric emptying in male HSD rats with EC50 = 0.12 and 0.29 .mu.g. Exendin-4 showed EC50 = 0.27 .mu.g under the same conditions.

IT 141758-74-9P, Exendin-4 (Heloderma suspectum)

203743-28-6P 203743-30-0P 203743-40-2P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of exendin peptide analogs as agonists for regulating gastrointestinal motility)

L2 ANSWER 33 OF 40 HCAPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 1997:577997 HCAPLUS  
 DOCUMENT NUMBER: 127:257827  
 TITLE: Novel signal transduction and peptide specificity of glucagon-like peptide receptor in 3T3-L1 adipocytes  
 AUTHOR(S): Montrose-Rafizadeh, Chahrzad; Yang, Huan; Wang, Yihong; Roth, Jesse; Montrose, Marshall H.; Adams, Lisa G.  
 CORPORATE SOURCE: Laboratory of Clinical Physiology, Gerontology Research Center, National Institute on Aging, NIH, Baltimore, MD, USA  
 SOURCE: Journal of Cellular Physiology (1997), 172(3), 275-283  
 CODEN: JCLLAX; ISSN: 0021-9541

PUBLISHER: Wiley-Liss  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 AB Glucagon-like peptide-1 (7-36) amide (GLP-1), in addn. to its well known effect of enhancing glucose-mediated insulin release, has been shown to have insulinomimetic effects and to enhance insulin-mediated glucose uptake and lipid synthesis in 3T3-L1 adipocytes. To elucidate the mechanisms of GLP-1 action in these cells, the authors studied the signal transduction and peptide specificity of the GLP-1 response. In 3T3-L1 adipocytes, GLP-1 caused a decrease in intracellular cAMP levels which is the opposite to the response obsd. in pancreatic beta cells in response to the same peptide. In 3T3-L1 adipocytes, free intracellular calcium was not modified by GLP-1. Peptide specificity was examd. to help det. if a different GLP receptor isoform was expressed in 3T3-L1 adipocytes vs. beta cells. Peptides with partial homol. to GLP-1 such as GLP-2, GLP-1 (1-36), and glucagon all lowered cAMP levels in 3T3-L1 adipocytes. In addn., an antagonist of pancreatic GLP-1 receptor, exendin-4 (9-39), acted as an agonist to decrease cAMP levels in 3T3-L1 adipocytes as did exendin-4 (1-39), a known agonist for the pancreatic GLP-1 receptor. Binding studies using <sup>125</sup>I-GLP-1 also suggest that pancreatic GLP-1 receptor isoform is not responsible for the effect of GLP-1 and related peptides in 3T3-L1 adipocytes. Based on these results, the authors propose that the major form of the GLP receptor in 3T3-L1 adipocytes is functionally different from the pancreatic GLP-1 receptor.  
 IT **141758-74-9**, Exendin 4 (*Heloderma suspectum*)  
 RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
 (signal transduction and peptide specificity of glucagon-like peptide receptor in 3T3-L1 adipocytes)

L2 ANSWER 34 OF 40 HCAPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 1997:567059 HCAPLUS  
 DOCUMENT NUMBER: 127:257697  
 TITLE: High potency antagonists of the pancreatic glucagon-like peptide-1 receptor  
 AUTHOR(S): Montrose-Rafizadeh, Chahrzad; Yang, Huan; Rodgers, Buel D.; Beday, Alvie; Pritchette, Louella A.; Eng, John  
 CORPORATE SOURCE: Laboratory of Clinical Physiology, NIA, National Institutes of Health, Baltimore, MD, 21224, USA  
 SOURCE: Journal of Biological Chemistry (1997), 272(34), 21201-21206  
 CODEN: JBCHA3; ISSN: 0021-9258  
 PUBLISHER: American Society for Biochemistry and Molecular Biology  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 AB GLP-1-(7-36)-amide and exendin-4-(1-39) are glucagon-like peptide-1 (GLP-1) receptor agonists, whereas exendin-(9-39) is the only known antagonist. To analyze the transition from agonist to antagonist and to identify the amino acid residues involved in ligand activation of the GLP-1 receptor, we used exendin analogs with successive N-terminal truncations. Chinese hamster ovary cells stably transfected with the rat GLP-1 receptor were assayed for changes in intracellular cAMP caused by the test peptides in the

absence or presence of half-maximal stimulatory doses of GLP-1. N-terminal truncation of a single amino acid reduced the agonist activity of the exendin peptide, whereas N-terminal truncation of 3-7 amino acids produced antagonists that were 4-10-fold more potent than exendin-(9-39). N-terminal truncation of GLP-1 by 2 amino acids resulted in weak agonist activity, but an 8-amino acid N-terminal truncation inactivated the peptide. Binding studies performed using  $^{125}\text{I}$ -labeled GLP-1 confirmed that all bioactive peptides specifically displaced tracer with high potency. In a set of exendin/GLP-1 chimeric peptides, substitution of GLP-1 sequences into exendin-(3-39) produced loss of antagonist activity with conversion to a weak agonist. The results show that receptor binding and activation occur in sep. domains of exendin, but they are more closely coupled in GLP-1.

IT **141758-74-9**, Exendin 4 (*Heloderma suspectum*)  
 RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study); PROC (Process)  
 (glucagon-like peptide-1 receptor high potency antagonists and structure-activity relations thereof)

L2 ANSWER 35 OF 40 HCPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 1997:127672 HCPLUS  
 DOCUMENT NUMBER: 126:223096  
 TITLE: Tissue-specific expression of unique mRNAs that encode proglucagon-derived peptides or exendin 4 in the lizard  
 AUTHOR(S): Chen, Yuqing E.; Drucker, Daniel J.  
 CORPORATE SOURCE: Toronto Hosp., Univ. Toronto, Toronto, ON, M5G 2C4, Can.  
 SOURCE: Journal of Biological Chemistry (1997), 272(7), 4108-4115  
 CODEN: JBCHA3; ISSN: 0021-9258  
 PUBLISHER: American Society for Biochemistry and Molecular Biology  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 AB Glucagon-like peptide 1 stimulates insulin secretion and inhibits glucagon secretion, gastric emptying, and feeding, suggesting it may be biol. useful for the treatment of diabetes. A lizard glucagon-like peptide 1(GLP-1)-related peptide, exendin 4, binds to the GLP-1 receptor and mimics the actions of GLP-1 in vivo. To det. the genetic relationship between exendin 4 and GLP-1, the authors analyzed the structure and expression of pancreatic and intestinal proglucagon mRNAs in the reptile *Heloderma suspectum*. Two different proglucagon cDNAs (lizard proglucagon I (LPI) and lizard proglucagon II (LPII)), with unique 3'-untranslated regions were identified. Two LPI mRNA transcripts, .apprx.1.6 and 2.1 kilobases, encoded glucagon and GLP-1 but not GLP-2 and were restricted in expression to the pancreas. In contrast, a 1.1-kilobase LPII mRNA transcript, encoding glucagon, GLP-1, and GLP-2 utilized a different 3'-untranslated region and was expressed in both pancreas and intestine. Lizard proglucagon mRNA transcripts were not detectable by reverse transcription-polymerase chain reaction or Northern blotting in salivary gland. A single class of lizard salivary gland proexendin cDNAs encoded the sequence of exendin 4 and a 45-amino acid exendin N-terminal peptide. Exendin mRNA transcripts were expressed in the salivary gland, but not pancreas or intestine.

09/756690

These data demonstrate that GLP-1 and exendin-4 represent related yet distinct peptide encoded by different genes in the lizard.

IT 188265-76-1, Exendin 4, pro- (Heloderma suspectum)

RL: PRP (Properties)

(amino acid sequence; unique mRNAs that encode proglucagon-derived peptides or exendin 4 tissue-specific expression in lizard)

L2 ANSWER 36 OF 40 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1995:675100 HCAPLUS

DOCUMENT NUMBER: 123:74913

TITLE: Exendin-3 and exendin-4 polypeptides, and pharmaceutical compositions comprising them

INVENTOR(S): Eng, John

PATENT ASSIGNEE(S): USA

SOURCE: U.S., 17 pp.

CODEN: USXXAM

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 5424286	A	19950613	US 1993-66480	19930524

PRIORITY APPLN. INFO.: US 1993-66480 19930524

AB This invention encompasses pharmaceutical compns. contg. exendin-3 or exendin-4, fragments thereof, or any combination thereof, and methods for the treatment of diabetes mellitus and the prevention of hyperglycemia.

IT 141758-74-9, Exendin 4 (Heloderma suspectum)

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(exendin-3 and exendin-4 polypeptides, and pharmaceutical compns. comprising them)

L2 ANSWER 37 OF 40 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1994:622490 HCAPLUS

DOCUMENT NUMBER: 121:222490

TITLE: Use of  $^{125}\text{I}$ -[Y39]exendin-4 to characterize exendin receptors on dispersed pancreatic acini and gastric chief cells from guinea pig

AUTHOR(S): Singh, Gurcharn; Eng, John; Raufman, Jean-Pierre  
CORPORATE SOURCE: Gastrointestinal Cell Biology Laboratory, State University of New York-Health Science Center at Brooklyn, 450 Clarkson Avenue-Box 1196, Brooklyn, NY, 11203-2098, USA

SOURCE: Regulatory Peptides (1994), 53(1), 47-59  
CODEN: REPPDY; ISSN: 0167-0115

DOCUMENT TYPE: Journal

LANGUAGE: English

AB We synthesized and iodinated an exendin-4 analog, [Y39]exendin-4 (700 Ci/mmol), for use as a radioligand to characterize exendin receptors on dispersed pancreatic acini and gastric chief cells from guinea pig. Binding of this bioactive radioligand was rapid, temp.-dependent and specific (not inhibited by other pancreatic or gastric secretagogues). Measurement of the ability of exendin-4 to

inhibit the binding of  $^{125}\text{I}$ -[Y39]exendin-4 indicated the presence of two classes of receptors. Pancreatic acini had 12.5 times 1010 binding sites/mg acinar protein of which 6% were high affinity ( $K_d = 0.5 \text{ nM}$ ) and 94% were low affinity ( $K_d = 0.1 \text{ }\mu\text{M}$ ). Chief cells had 3370 binding sites/cell of which 9% were high affinity ( $K_d = 0.3 \text{ nM}$ ) and 91% were low affinity ( $K_d = 0.2 \text{ }\mu\text{M}$ ). Washing with 0.2 M acetic acid (pH 2.5), 0.2 M glycine (pH 10.5), or trypsin (100  $\mu\text{g/mL}$ ) after 30 min incubation at 37 degree., indicated that 63 and 49% of radioligand was internalized in acini and chief cells, resp. Truncated glucagon-like peptide-1 (tGLP-1), a mammalian peptide sharing 53% homol. with exendin-4, inhibited radioligand binding at the same concns. that altered secretion from acini and chief cells. Glucagon, GLP-1 and GLP-2 inhibited  $^{125}\text{I}$ -[Y39]exendin-4 binding only at concns.  $\geq 100 \text{ nM}$ . Exendin(9-39)NH<sub>2</sub>, a specific exendin-receptor antagonist, potently inhibited  $^{125}\text{I}$ -[Y39] exendin-4 binding ( $IC_{50} = 6.1$  and  $3.5 \text{ nM}$  in acini and chief cells, resp.). In pancreatic acini and gastric chief cells from guinea pig, exendin-3, exendin-4 and tGLP-1 increase cellular cAMP and modulate enzyme secretion by interacting with high-affinity exendin receptors.  $^{125}\text{I}$ -[Y39] exendin-4 is a useful radioligand for studying exendin receptors.

IT

**141758-74-9**

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)  
(cAMP formation and enzyme secretion by pancreas acinus and stomach chief cells response to)

L2 ANSWER 38 OF 40 HCAPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 1993:597526 HCAPLUS  
 DOCUMENT NUMBER: 119:197526  
 TITLE: Exendin-4 is a high potency agonist and truncated exendin-(9-39)-amide an antagonist at the glucagon-like peptide 1-(7-36)-amide receptor of insulin-secreting .beta.-cells  
 AUTHOR(S): Goeke, Ruediger; Fehmann, Hans Christoph; Linn, Thomas; Schmidt, Harald; Krause, Michael; Eng, John; Goeke, Burkhard  
 CORPORATE SOURCE: Dep. Intern. Med., Philipps Univ., Marburg, 3550, Germany  
 SOURCE: Journal of Biological Chemistry (1993), 268(26), 19650-5  
 DOCUMENT TYPE: CODEN: JBCHA3; ISSN: 0021-9258  
 LANGUAGE: English  
 AB Exendin-4 purified from *Heloderma suspectum* venom shows structural relationship to the important incretin hormone glucagon-like peptide 1-(7-36)-amide (GLP-1). The authors demonstrate that exendin-4 and truncated exendin-(9-39)-amide specifically interact with the GLP-1 receptor on insulinoma-derived cells and on lung membranes. Exendin-4 displaced  $^{125}\text{I}$ -GLP-1, and unlabeled GLP-1 displaced  $^{125}\text{I}$ -exendin-4 from the binding site at rat insulinoma-derived RINm5F cells. Exendin-4 had, like GLP-1, a pronounced effect on intracellular cAMP generation, which was reduced by exendin-(9-39)-amide. When combined, GLP-1 and exendin-4 showed additive action on cAMP. They each competed with the radiolabeled version of the other peptide in crosslinking expts. The apparent mol. mass of the resp. ligand-binding protein complex was 63,000 Da. Exendin-(9-39)-amide abolished the crosslinking of both peptides.

Exendin-4, like GLP-1, stimulated dose dependently the glucose-induced insulin secretion in isolated rat islets, and, in mouse insulinoma .beta.TC-1 cells, both peptides stimulated the proinsulin gene expression at the level of transcription. Exendin-(9-39)-amide reduced these effects. In conclusion, exendin-4 is an agonist and exendin-(9-39)-amide is a specific GLP-1 receptor antagonist.

IT **141758-74-9**

RL: BIOL (Biological study)  
(glucagon-like peptide 1-(7-36)-amide receptor of .beta.-cells and lung response to)

L2 ANSWER 39 OF 40 HCAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 1992:564310 HCAPLUS  
DOCUMENT NUMBER: 117:164310  
TITLE: Truncated glucagon-like peptide-1 interacts with exendin receptors on dispersed acini from guinea pig pancreas. Identification of a mammalian analogue of the reptilian peptide exendin-4  
AUTHOR(S): Raufman, Jean Pierre; Singh, Latika; Singh, Gurcharn; Eng, John  
CORPORATE SOURCE: Health Sci. Cent., State Univ. New York, Brooklyn, NY, 11203-2098, USA  
SOURCE: Journal of Biological Chemistry (1992), 267(30), 21432-7  
CODEN: JBCHA3; ISSN: 0021-9258  
DOCUMENT TYPE: Journal  
LANGUAGE: English  
AB To find mammalian analogs of exendin-4, a peptide from Helodermatidae venoms that interacts with newly discovered exendin receptors on dispersed acini from guinea pig pancreas, the actions of glucagon-like peptide-1 [GLP-1(1-37)], its truncated form GLP-1(7-36)NH<sub>2</sub>, GLP-2(1-34), and pituitary adenylate cyclase-activating peptide were examined and compared with secretin, VIP, and glucagon. Only the truncated form of glucagon-like peptide-1, GLP-1(7-36)NH<sub>2</sub> mimicked the actions of exendin-4. Like exendin-4, GLP-1(7-36)NH<sub>2</sub> increased acinar cAMP without stimulating amylase release. GLP-1(7-36)NH<sub>2</sub>-induced increases in cAMP were inhibited progressively by increasing concns. of the specific exendin-receptor antagonist, exendin(9-39)NH<sub>2</sub>. In dispersed acini from guinea pig and rat pancreas, concns. of GLP-1(7-36)NH<sub>2</sub> that stimulated increases in cAMP caused potentiation of cholecystokinin-induced amylase release. Binding of <sup>125</sup>I-[Y39]exendin-4 or <sup>125</sup>I-GLP-1(7-36)NH<sub>2</sub> to dispersed acini from guinea pig pancreas was inhibited by adding increasing concns. of unlabeled exendin-4 or GLP-1(7-36)NH<sub>2</sub>. Thus, the mammalian peptide GLP-1(7-36)NH<sub>2</sub> interacts with exendin receptors on dispersed acini from guinea pig pancreas. Exendin(9-39)NH<sub>2</sub>, a competitive antagonist of the actions of GLP-1(7-36)NH<sub>2</sub> in pancreatic acini, may be a useful tool for examining the physiol. actions of this peptide.

IT **141758-74-9**

RL: BIOL (Biological study)  
(glucagon-like peptide 1 truncated form as mammalian analog of)

L2 ANSWER 40 OF 40 HCAPLUS COPYRIGHT 2003 ACS  
ACCESSION NUMBER: 1992:402472 HCAPLUS  
DOCUMENT NUMBER: 117:2472  
TITLE: Isolation and characterization of exendin-4, an

09/756690

AUTHOR(S): Eng, John; Kleinman, Wayne A.; Singh, Latika; Singh, Gurcharn; Raufman, Jean Pierre  
CORPORATE SOURCE: Solomon A Berson Res. Lab., Veterans Aff. Med. Cent., Bronx, NY, 10468, USA  
SOURCE: Journal of Biological Chemistry (1992), 267(11), 7402-5  
DOCUMENT TYPE: Journal  
LANGUAGE: English

AB An amino acid sequencing assay for peptides contg. an amino-terminal histidine residue (His1) was used to isolate a 39-amino acid peptide, exendin-4, from *H. suspectum* venom. Exendin-4 differs from exendin-3 by two amino acid substitutions, Gly2-Glu3 in place of Ser2-Asp3, but is otherwise identical. The structural differences make exendin-4 distinct from exendin-3 in its bioactivity. In dispersed acini from guinea pig pancreas, natural and synthetic exendin-4 stimulate a monophasic increase in cAMP beginning at 100 pM that plateaus at 10 nM. The exendin-4-induced increase in cAMP is inhibited progressively by increasing concns. of the exendin receptor antagonist, exendin-(9-39) amide. Unlike exendin-3, exendin-4 does not stimulate a second rise in acinar cAMP at concns. >100 nM, does not stimulate amylase release, and does not inhibit the binding of radiolabeled vasoactive intestinal peptide to acini. This indicates that in dispersed pancreatic acini, exendin-4 interacts only with the recently described exendin receptor.

IT **141758-74-9**

RL: PRP (Properties)  
(amino acid sequence of, complete)

=> sel hit 12 1-40 rn  
E72 THROUGH E101 ASSIGNED

FILE 'REGISTRY' ENTERED AT 10:32:52 ON 14 FEB 2003

L3 29 SEA FILE=REGISTRY ABB=ON PLU=ON (141758-74-9/BI OR  
203743-40-2/BI OR 188265-76-1/BI OR 210829-56-4/BI OR  
210829-59-7/BI OR 335149-21-8/BI OR 203743-28-6/BI OR  
203743-30-0/BI OR 211430-62-5/BI OR 211430-73-8/BI OR  
305814-59-9/BI OR 305815-28-5/BI OR 305815-30-9/BI OR  
305818-90-0/BI OR 308244-92-0/BI OR 308245-14-9/BI OR  
308245-46-7/BI OR 308815-99-8/BI OR 309257-15-6/BI OR  
309728-25-4/BI OR 309729-73-5/BI OR 309729-78-0/BI OR  
320367-11-1/BI OR 320367-31-5/BI OR 435950-95-1/BI OR  
437124-38-4/BI OR 437124-39-5/BI OR 437124-51-1/BI OR  
437124-53-3/BI OR 474444-81-0/BI)

L4 29 L3 AND L1

L4 ANSWER 1 OF 29 REGISTRY COPYRIGHT 2003 ACS

RN **474444-81-0** REGISTRY

CN L-Serine, L-histidylglycyl-L-.alpha.-glutamylglycyl-L-threonyl-L-phenylalanyl-L-threonyl-L-seryl-L-.alpha.-aspartyl-L-leucyl-L-seryl-L-lysyl-L-glutaminyl-L-methionyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-alanyl-L-valyl-L-arginyl-L-leucyl-L-phenylalanyl-L-isoleucyl-L-.alpha.-glutamyl-L-tryptophyl-L-leucyl-L-lysyl-L-asparaginylglycylglycyl-L-prolyl-L-seryl-L-serylglycyl-L-

Searcher : Shears 308-4994

09/756690

alanyl-L-prolyl-L-prolyl-L-prolyl- (9CI) (CA INDEX NAME)  
OTHER NAMES:

CN 5: PN: WO02085406 SEQID: 9 unclaimed protein  
CI MAN  
SQL 39

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPPS  
===== ===== ===== =====

HITS AT: 1-39

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

REFERENCE 1: 137:346934

L4 ANSWER 2 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN 437124-53-3 REGISTRY  
CN exendin-4 fusion protein with a linker and human IgG1 fragment (9CI)  
(CA INDEX NAME)  
OTHER NAMES:

CN 29: PN: WO0246227 SEQID: 31 claimed protein  
CI MAN  
SQL 287

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPSG GGGSGGGGSG  
===== ===== ===== =====  
51 GGGSAEPKSC DKHTCPPCP APELLGGPSV FLFPPKPKDT LMISRTPEVT  
101 CVVVDVSHED PEVKFNWYVD GVEVHNNAKTK PREEQYNSTY RVVSVLTVLH  
151 QDWLNGKEYK CKVSNKALPA PIEKTISKAK GQPREPQVYT LPPSREEMTK  
201 NQVSLTCLVK GFYPSDIAVE WESNGQPENN YKTTPPVLDS DGSSFFLYSKL  
251 TVDKSRWQQG NVFSCSVMHE ALHNHYTQKS LSLSPGK

HITS AT: 1-39

REFERENCE 1: 137:28591

L4 ANSWER 3 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN 437124-51-1 REGISTRY  
CN exendin-4 fusion protein with human IgG1 fragment (9CI) (CA INDEX  
NAME)  
OTHER NAMES:

CN 27: PN: WO0246227 SEQID: 29 claimed protein  
CI MAN  
SQL 272

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPSA EPKSCDKTHT  
===== ===== ===== =====  
51 CPPCPAPELL GGPSVFLFPP KPKDTLMISR TPEVTCVVVD VSHEDPEVKF  
101 NWYVDGVEVH NAKTKPREQE YNSTYRVVSV LTVLHQDWLN GKEYKCKVSN  
151 KALPAPIEKT ISKAKGQPREG PVQYTLPPSR EEMTKNQVSL TCLVKGFYPS  
201 DIAVEWESNG QPENNYKTTP PVLDSDGSFF LYSKLTVDKS RWQQGNVFSC  
251 SVMHEALHNN YTQKSLSLSP GK

HITS AT: 1-39

REFERENCE 1: 137:28591

L4 ANSWER 4 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN 437124-39-5 REGISTRY  
CN exendin-4 fusion protein with linker and human serum albumin (9CI)  
(CA INDEX NAME)

09/756690

OTHER NAMES:

CN 16: PN: WO0246227 SEQID: 17 claimed protein  
CI MAN  
SQL 640

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPPSG GGGGSGGGGS  
===== ===== ===== =====  
51 GGGGSDAHKS EVAHFKDLG EENFKALVLI AFAQYLQQCP FEDHVKLVNE  
101 VTEFAKTCVA DESAENCDKS LHTLFGDKLC TVATLRETYG EMADCCAKQE  
151 PERNECFLQH KDDNPNLPRL VRPEVDVMCT AFHDNEETFL KYLYEIARR  
201 HPFYAPELL FFAKRYKAAC TECCQAADKA ACLLPKLDEL RDEGKASSAK  
251 QRLKCASLQK FGERAFKAWA VARLSQRFPK AEFAEVSKLV TDLTKVHTEC  
301 CHGDLLECAD DRADLAKYIC ENQDSISSLK KECCEKPLLE KSHCIAEVEN  
351 DEMPADLPSL AADFVESKDV CKNYAEAKDV FLGMFLYEA RRHPDYSVVL  
401 LLRLAKTYET TLEKCCAAAD PHECYAKVFD EFKPLVEEPQ NLIKQNCELF  
451 EQLGEYKFQN ALLVRYTKV PQVSTPTLVE VSRNLGKVGK KCCKHPEAKR  
501 MPCADEYLSV VLNQLCVLHE KTPVSDRVTK CCTESLVNRR PCFSALEVDE  
551 TYVPKEFNAE TFTFHADICT LSEKERQIKK QTALVELVKH KPKATKQQLK  
601 AVMDDDFAAFV EKCCKADDKE TCFAEKGKLV ASQAAALGL

HITS AT: 1-39

REFERENCE 1: 137:28591

L4 ANSWER 5 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN 437124-38-4 REGISTRY  
CN exendin-4 fusion protein with human serum albumin (9CI) (CA INDEX  
NAME)  
OTHER NAMES:  
CN 15: PN: WO0246227 SEQID: 16 claimed protein  
CI MAN  
SQL 624

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPPSD AHKSEVAHRF  
===== ===== ===== =====  
51 KDLGEENFKA LVLIAFAQYL QQCPFEDHVK LVNEVTÉFAK TCVADESAEN  
101 CDKSLHTLFG DKLCTVATLR ETYGEMADCC AKQEPERNEC FLQHKDDNPN  
151 LPRLVRPEVD VMCTAFHDNE ETFLKKYLYE IARRHPYFA PELLFFAKRY  
201 KAAFTCCQA ADKAACLLPK LDELRDEGKA SSAKQLKCA SLQKFGERA  
251 KAWAVARLSQ RFPKAEFAEV SKLVTDLTKV HTECCHGDLL ECADDRADLA  
301 KYICENQDSI SSKLKECCEK PLLEKSHCIA EVENDEMPAD LPSLAADFVE  
351 SKDVCKNYAE AKDVFLGMFL YEYARRHPDY SVVLLRLAK TYETTLEKCC  
401 AAADPHCYA KVFDEFKPLV EEPQNLIKQN CELFEQLGEY KFQNALLVRY  
451 TKKVPQVSTP TLVEVSRNLG KVGSCKCKHP EAKRMPCAED YLSVVLNQLC  
501 VLHEKTPVSD RVTKCCTESL VNRRPCFSAL EVDETYVPKE FNAETFTFHA  
551 DICTLSEKER QIKKQTALVE LVKHKPKATK EQLKAVMDDF AAFVEKCKA  
601 DDKETCFTEE GKVLVAASQA ALGL

HITS AT: 1-39

REFERENCE 1: 137:28591

L4 ANSWER 6 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN 435950-95-1 REGISTRY  
CN L-Serinamide, L-histidylglycyl-L-.alpha.-glutamylglycyl-L-threonyl-L-phenylalanyl-L-threonyl-L-seryl-L-.alpha.-aspartyl-L-leucyl-L-seryl-L-lysyl-L-glutaminyl-L-methionyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-alanyl-L-valyl-L-arginyl-L-leucyl-L-phenylalanyl-L-isoleucyl-L-.alpha.-glutamyl-L-tryptophyl-L-leucyl-L-lysyl-L-asparaginylglycylglycyl-L-prolyl-L-seryl-L-serylglycyl-L-

09/756690

CI alanyl-L-prolyl-L-prolyl-L-prolyl- (9CI) (CA INDEX NAME)  
MAN  
SQL 39

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPPS  
===== ===== ===== ===== =====

HITS AT: 1-39

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

REFERENCE 1: 137:28591

L4 ANSWER 7 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN 320367-31-5 REGISTRY  
CN L-Lysinamide, L-histidylglycyl-L-.alpha.-glutamylglycyl-L-threonyl-L-phenylalanyl-L-threonyl-L-seryl-L-.alpha.-aspartyl-L-leucyl-L-seryl-L-lysyl-L-glutaminyl-L-methionyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-alanyl-L-valyl-L-arginyl-L-leucyl-L-phenylalanyl-L-isoleucyl-L-.alpha.-glutamyl-L-tryptophyl-L-leucyl-L-lysyl-L-asparaginylglycylglycyl-L-prolyl-L-seryl-L-serylglycyl-L-alanyl-L-prolyl-L-prolyl-L-prolyl-L-seryl-N6-(1-oxohexadecyl)-L-lysyl-L-lysyl-L-lysyl-L-lysyl-L-lysyl- (9CI) (CA INDEX NAME)

CI MAN  
SQL 46

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPPSK KKKKK  
===== ===== ===== ===== =====

HITS AT: 1-39

REFERENCE 1: 134:110470

L4 ANSWER 8 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN 320367-11-1 REGISTRY  
CN L-Lysinamide, L-histidylglycyl-L-.alpha.-glutamylglycyl-L-threonyl-L-phenylalanyl-L-threonyl-L-seryl-L-.alpha.-aspartyl-L-leucyl-L-seryl-L-lysyl-L-glutaminyl-L-methionyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-alanyl-L-valyl-L-arginyl-L-leucyl-L-phenylalanyl-L-isoleucyl-L-.alpha.-glutamyl-L-tryptophyl-L-leucyl-L-lysyl-L-asparaginylglycylglycyl-L-prolyl-L-seryl-L-serylglycyl-L-alanyl-L-prolyl-L-prolyl-L-prolyl-L-seryl-L-lysyl-L-lysyl-L-lysyl- (9CI) (CA INDEX NAME)

CI MAN  
SQL 45

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPPSK KKKKK  
===== ===== ===== ===== =====

HITS AT: 1-39

REFERENCE 1: 134:110470

L4 ANSWER 9 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN 309729-78-0 REGISTRY  
CN Exendin 4 (Heloderma suspectum), 39a-[N6-[21-(2,5-dihydro-2,5-dioxo-1H-pyrrol-1-yl)-1,10,19-trioxa-3,6,12,15-tetraoxa-9,18-diazaheneicos-1-yl]-L-lysinamide]-, pentakis(trifluoroacetate) (salt) (9CI) (CA INDEX NAME)

OTHER NAMES:

09/756690

CN 34: PN: WO0069911 PAGE: 66 claimed sequence  
SQL 40

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPSK  
===== ===== ===== ===== =====  
HITS AT: 1-39

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPSK  
===== ===== ===== ===== =====  
HITS AT: 1-39

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPSK  
===== ===== ===== ===== =====  
HITS AT: 1-39

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

REFERENCE 1: 134:13338

L4 ANSWER 10 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN 309729-73-5 REGISTRY  
CN Exendin 4 (Heloderma suspectum), 39a-[N6-[21-(2,5-dihydro-2,5-dioxo-1H-pyrrol-1-yl)-1,10,19-trioxa-3,6,12,15-tetraoxa-9,18-diazaheneicos-1-yl]-L-lysinamide]- (9CI) (CA INDEX NAME)  
CI COM, MAN  
SQL 40

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPSK  
===== ===== ===== ===== =====  
HITS AT: 1-39

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

REFERENCE 1: 134:13338

L4 ANSWER 11 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN 309728-25-4 REGISTRY  
CN L-Lysinamide, L-histidylglycyl-L-.alpha.-glutamylglycyl-L-threonyl-L-phenylalanyl-L-threonyl-L-seryl-L-.alpha.-aspartyl-L-leucyl-L-seryl-L-lysyl-L-glutaminyl-L-methionyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-alanyl-L-valyl-L-arginyl-L-leucyl-L-phenylalanyl-L-isoleucyl-L-.alpha.-glutamyl-L-tryptophyl-L-leucyl-L-lysyl-L-asparaginylglycylglycyl-L-prolyl-L-seryl-L-serylglycyl-L-alanyl-L-prolyl-L-prolyl-L-seryl-L-seryl- (9CI) (CA INDEX NAME)  
OTHER NAMES:

CN 33: PN: WO0069911 PAGE: 63 claimed sequence  
CI MAN  
SQL 40

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPSK  
===== ===== ===== ===== =====  
HITS AT: 1-39

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

09/756690

REFERENCE 1: 134:13338

L4 ANSWER 12 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN 309257-15-6 REGISTRY  
CN 192: PN: WO0069900 SEQID: 371 unclaimed protein (9CI) (CA INDEX  
NAME)  
CI MAN  
SQL 40

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPSK  
===== ===== ===== =====

HITS AT: 1-39

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

REFERENCE 1: 134:21425

L4 ANSWER 13 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN 308815-99-8 REGISTRY  
CN L-Lysine, L-histidylglycyl-L-.alpha.-glutamylglycyl-L-threonyl-L-phenylalanyl-L-threonyl-L-seryl-L-.alpha.-aspartyl-L-leucyl-L-seryl-L-lysyl-L-glutaminyl-L-methionyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-alanyl-L-valyl-L-arginyl-L-leucyl-L-phenylalanyl-L-isoleucyl-L-.alpha.-glutamyl-L-tryptophyl-L-leucyl-L-lysyl-L-asparaginylglycylglycyl-L-prolyl-L-seryl-L-serylglycyl-L-alanyl-L-prolyl-L-prolyl-L-prolyl-L-seryl- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 18: PN: WO0069911 SEQID: 18 claimed protein  
CI MAN  
SQL 40

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPSK  
===== ===== ===== =====

HITS AT: 1-39

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

REFERENCE 1: 134:13338

L4 ANSWER 14 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN 308245-46-7 REGISTRY  
CN Exendin 4 (Heloderma suspectum), 39a-[N6-[15-(2,5-dihydro-2,5-dioxo-1H-pyrrol-1-yl)-1,7,13-trioxa-3,9-dioxa-6,12-diazapentadec-1-yl]-L-lysinamide]-, pentakis(trifluoroacetate) (salt) (9CI) (CA INDEX NAME)  
SQL 40

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPSK  
===== ===== ===== =====

HITS AT: 1-39

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPSK  
===== ===== ===== =====

HITS AT: 1-39

09/756690

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

SEQ 1 HGEGTFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPSK  
===== ===== ===== ===== =====  
HITS AT: 1-39

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

REFERENCE 1: 134:21425

L4 ANSWER 15 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN 308245-14-9 REGISTRY  
CN Exendin 4 (Heloderma suspectum), 39a-[N6-[3-(2,5-dihydro-2,5-dioxo-1H-pyrrol-1-yl)-1-oxopropyl]-L-lysinamide]-, pentakis(trifluoroacetate) (salt) (9CI) (CA INDEX NAME)  
SQL 40

SEQ 1 HGEGTFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPSK  
===== ===== ===== ===== =====  
HITS AT: 1-39

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

SEQ 1 HGEGTFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPSK  
===== ===== ===== ===== =====  
HITS AT: 1-39

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

SEQ 1 HGEGTFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPSK  
===== ===== ===== ===== =====  
HITS AT: 1-39

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

REFERENCE 1: 134:21425

L4 ANSWER 16 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN 308244-92-0 REGISTRY  
CN Exendin 4 (Heloderma suspectum), 39a-[N6-[3-(2,5-dihydro-2,5-dioxo-1H-pyrrol-1-yl)-1-oxopropyl]-L-lysinamide]- (9CI) (CA INDEX NAME)  
OTHER NAMES:  
CN 32: PN: WO0069911 PAGE: 62 claimed sequence  
CI COM, MAN  
SQL 40

SEQ 1 HGEGTFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPSK  
===== ===== ===== ===== =====  
HITS AT: 1-39

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

REFERENCE 1: 134:13338

L4 ANSWER 17 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN 305818-90-0 REGISTRY  
CN Poly(oxy-1,2-ethanediyl), .alpha.-hydro-.omega.-hydroxy-, 27-ether with L-histidylglycyl-L-.alpha.-glutamylglycyl-L-threonyl-L-

09/756690

phenylalanyl-L-threonyl-L-seryl-L-.alpha.-aspartyl-L-leucyl-L-seryl-L-lysyl-L-glutaminyl-L-methionyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-alanyl-L-valyl-L-arginyl-L-leucyl-L-phenylalanyl-L-isoleucyl-L-.alpha.-glutamyl-L-tryptophyl-L-leucyl-N6-(2-hydroxyethyl)-L-lysyl-L-asparaginylglycylglycyl-L-prolyl-L-seryl-L-serylglycyl-L-alanyl-L-prolyl-L-prolyl-L-prolyl-L-serinamide (9CI) (CA INDEX NAME)

CI PMS, MAN  
SQL 39

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPPS  
===== ===== ===== =====

HITS AT: 1-39

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

REFERENCE 1: 133:359242

L4 ANSWER 18 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN 305815-30-9 REGISTRY  
CN L-Serinamide, N-ethyl-L-histidylglycyl-L-.alpha.-glutamylglycyl-L-threonyl-L-phenylalanyl-L-threonyl-L-seryl-L-.alpha.-aspartyl-L-leucyl-L-seryl-L-lysyl-L-glutaminyl-L-methionyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-alanyl-L-valyl-L-arginyl-L-leucyl-L-phenylalanyl-L-isoleucyl-L-.alpha.-glutamyl-L-tryptophyl-L-leucyl-L-lysyl-L-asparaginylglycylglycyl-L-prolyl-L-seryl-L-serylglycyl-L-alanyl-L-prolyl-L-prolyl-L-prolyl- (9CI) (CA INDEX NAME)

CI MAN  
SQL 39

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPPS  
===== ===== ===== =====

HITS AT: 1-39

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

REFERENCE 1: 133:359242

L4 ANSWER 19 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN 305815-28-5 REGISTRY  
CN L-Serinamide, N-acetyl-L-histidylglycyl-L-.alpha.-glutamylglycyl-L-threonyl-L-phenylalanyl-L-threonyl-L-seryl-L-.alpha.-aspartyl-L-leucyl-L-seryl-L-lysyl-L-glutaminyl-L-methionyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-alanyl-L-valyl-L-arginyl-L-leucyl-L-phenylalanyl-L-isoleucyl-L-.alpha.-glutamyl-L-tryptophyl-L-leucyl-L-lysyl-L-asparaginylglycylglycyl-L-prolyl-L-seryl-L-serylglycyl-L-alanyl-L-prolyl-L-prolyl-L-prolyl- (9CI) (CA INDEX NAME)

CI MAN  
SQL 39

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPPS  
===== ===== ===== =====

HITS AT: 1-39

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

09/756690

REFERENCE 1: 133:359242

L4 ANSWER 20 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN **305814-59-9** REGISTRY  
CN Poly(oxy-1,2-ethanediyl), .alpha.-hydro-.omega.-hydroxy-, 12-ether  
with L-histidylglycyl-L-.alpha.-glutamylglycyl-L-threonyl-L-  
phenylalanyl-L-threonyl-L-seryl-L-.alpha.-aspartyl-L-leucyl-L-seryl-  
N6-(2-hydroxyethyl)-L-lysyl-L-glutaminyl-L-methionyl-L-.alpha.-  
glutamyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-alanyl-L-valyl-L-  
arginyl-L-leucyl-L-phenylalanyl-L-isoleucyl-L-.alpha.-glutamyl-L-  
tryptophyl-L-leucyl-L-lysyl-L-asparaginylglycylglycyl-L-prolyl-L-  
seryl-L-serylglycyl-L-alanyl-L-prolyl-L-prolyl-L-prolyl-L-serinamide  
(9CI) (CA INDEX NAME)  
CI PMS, MAN  
SQL 39

SEQ 1 HGEGTFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPPS  
===== ===== ===== =====

HITS AT: 1-39

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

REFERENCE 1: 133:359242

L4 ANSWER 21 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN **211430-73-8** REGISTRY  
CN Exendin ENTP (Heloderma horridum) (9CI) (CA INDEX NAME)  
CI MAN  
SQL 64

SEQ 1 MPVESGLSSE DSASSESFAS KIKRHGEGLTF TSDLSKQME EAVRLFIEWL  
===== ===== =====

51 KNGGPSSGAP PPSG  
===== ==

HITS AT: 25-63

REFERENCE 1: 129:185369

L4 ANSWER 22 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN **211430-62-5** REGISTRY  
CN Exendin 4 (Heloderma suspectum), 39a-glycine- (9CI) (CA INDEX NAME)  
OTHER NAMES:  
CN 48-87-Exendin ENTP (Heloderma horridum)  
CI MAN  
SQL 40

SEQ 1 HGEGTFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPPSG  
===== ===== ===== =====

HITS AT: 1-39

REFERENCE 1: 129:185369

L4 ANSWER 23 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN **210829-59-7** REGISTRY  
CN Exendin 4 (Heloderma suspectum), 36-[(4R)-4-thiazolidinecarboxylic  
acid]-37-[(4R)-4-thiazolidinecarboxylic acid]-38-[(4R)-4-  
thiazolidinecarboxylic acid]- (9CI) (CA INDEX NAME)  
OTHER NAMES:

09/756690

CN 22: PN: WO0073331 FIGURE: 1 claimed sequence  
CI MAN  
SQL 39

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPPS  
===== ===== ===== ===== =====  
HITS AT: 1-39

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

REFERENCE 1: 134:37033

REFERENCE 2: 129:149256

L4 ANSWER 24 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN **210829-56-4** REGISTRY  
CN L-Serinamide, L-histidylglycyl-L-.alpha.-glutamylglycyl-L-threonyl-L-phenylalanyl-L-threonyl-L-seryl-L-.alpha.-aspartyl-L-leucyl-L-seryl-L-lysyl-L-glutaminyl-L-methionyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-alanyl-L-valyl-L-arginyl-L-leucyl-L-phenylalanyl-L-isoleucyl-L-.alpha.-glutamyl-L-tryptophyl-L-leucyl-L-lysyl-L-asparaginylglycylglycyl-(4R)-4-thiazolidinecarbonyl-L-seryl-L-serylglycyl-L-alanyl-(4R)-4-thiazolidinecarbonyl-(4R)-4-thiazolidinecarbonyl-(4R)-4-thiazolidinecarbonyl- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 21: PN: WO0073331 FIGURE: 1 claimed sequence  
CI MAN  
SQL 39

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPPS  
===== ===== ===== ===== =====  
HITS AT: 1-39

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

REFERENCE 1: 134:37033

REFERENCE 2: 129:149256

L4 ANSWER 25 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN **203743-40-2** REGISTRY  
CN Exendin 4 (Heloderma suspectum), 39-L-serine- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 12: PN: WO0069911 SEQID: 12 claimed protein  
CN 170: PN: WO0069900 SEQID: 349 unclaimed protein  
CN 1: PN: WO0009666 SEQID: 9 unclaimed protein  
CN 2: PN: WO0151078 SEQID: 2 unclaimed protein  
CN 48-86-Exendin ENTP (Heloderma horridum)  
CN 4: PN: US6284725 SEQID: 9 unclaimed protein  
CN 4: PN: WO0066138 PAGE: 13 unclaimed protein  
CN 4: PN: WO0066142 TABLE: 1 unclaimed protein  
CN 4: PN: WO0123420 PAGE: 6 unclaimed protein  
CN 8: PN: WO0077039 TABLE: 1 unclaimed protein  
CI MAN  
SQL 39

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPPS

09/756690

HITS AT: 1-39

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

REFERENCE 1: 135:205920  
REFERENCE 2: 135:132445  
REFERENCE 3: 134:290749  
REFERENCE 4: 134:51920  
REFERENCE 5: 134:37033  
REFERENCE 6: 134:21425  
REFERENCE 7: 134:13338  
REFERENCE 8: 133:345167  
REFERENCE 9: 133:345166  
REFERENCE 10: 132:175839

L4 ANSWER 26 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN **203743-30-0** REGISTRY  
CN Exendin 4 (*Heloderma suspectum*), 36-(4-thiazolidinecarboxylic acid)-37-(4-thiazolidinecarboxylic acid)-38-(4-thiazolidinecarboxylic acid)- (9CI) (CA INDEX NAME)  
CI MAN  
SQL 39

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPPS

HITS AT: 1-39

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

REFERENCE 1: 128:192936

L4 ANSWER 27 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN **203743-28-6** REGISTRY  
CN L-Serinamide, L-histidylglycyl-L-.alpha.-glutamylglycyl-L-threonyl-L-phenylalanyl-L-threonyl-L-seryl-L-.alpha.-aspartyl-L-leucyl-L-seryl-L-lysyl-L-glutaminyl-L-methionyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-alanyl-L-valyl-L-arginyl-L-leucyl-L-phenylalanyl-L-soleucyl-L-.alpha.-glutamyl-L-tryptophyl-L-leucyl-L-lysyl-L-asparaginylglycylglycyl-4-thiazolidinecarbonyl-L-seryl-L-serylglycyl-L-alanyl-4-thiazolidinecarbonyl-4-thiazolidinecarbonyl-4-thiazolidinecarbonyl- (9CI) (CA INDEX NAME)  
CI MAN  
SQL 39

SEQ 1 HGETFTSDL SKQMEEEAVR LFIEWLKNGG PSSGAPPPS

HITS AT: 1-39

09/756690

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

REFERENCE 1: 128:192936

L4 ANSWER 28 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN 188265-76-1 REGISTRY  
CN Exendin 4, pro- (Heloderma suspectum) (9CI) (CA INDEX NAME)  
OTHER NAMES:  
CN Exendin 4 (Heloderma suspectum precursor)  
CN Exendin ENTP (Heloderma horridum pro-)  
CI MAN  
SQL 87

SEQ 1 MKIILWLWCVF GLFLATLFPI SWQMPVESGL SSEDSASSES FASKIKRHGE  
=====

51 GTFTSDLSKQ MEEEAVRLFI EWLKNGGPSS GAPPPSG  
=====

HITS AT: 48-86

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

REFERENCE 1: 137:346599

REFERENCE 2: 129:185369

REFERENCE 3: 129:78077

REFERENCE 4: 126:223096

L4 ANSWER 29 OF 29 REGISTRY COPYRIGHT 2003 ACS  
RN 141758-74-9 REGISTRY  
CN Exendin 4 (Heloderma suspectum) (9CI) (CA INDEX NAME)  
OTHER CA INDEX NAMES:  
CN Exendin 3 (Heloderma horridum), 2-glycine-3-L-glutamic acid-  
OTHER NAMES:  
CN 12: PN: WO0041546 FIGURE: 2 claimed protein  
CN 2: PN: WO0066629 FIGURE: 2 unclaimed sequence  
CN 3: PN: WO0041548 PAGE: 65 unclaimed protein  
CN AC 2993  
CN AC 2993A  
CN Exenatide  
CN Exendin-4 (Heloderma suspectum)  
CN L-Serinamide, L-histidylglycyl-L-.alpha.-glutamylglycyl-L-threonyl-L-phenylalanyl-L-threonyl-L-seryl-L-.alpha.-aspartyl-L-leucyl-L-seryl-L-lysyl-L-glutaminyl-L-methionyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-.alpha.-glutamyl-L-alanyl-L-valyl-L-arginyl-L-leucyl-L-phenylalanyl-L-isoleucyl-L-.alpha.-glutamyl-L-tryptophyl-L-leucyl-L-lysyl-L-asparaginylglycylglycyl-L-prolyl-L-seryl-L-serylglycyl-L-alanyl-L-prolyl-L-prolyl-L-prolyl-  
CI MAN  
SQL 39

SEQ 1 HGETFTSDL SKQMEAVR LFIEWLKNGG PSSGAPPPS  
=====

HITS AT: 1-39

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

09/756690

REFERENCE 1: 138:14048

REFERENCE 2: 137:311200

REFERENCE 3: 137:311199

REFERENCE 4: 137:247879

REFERENCE 5: 137:109267

REFERENCE 6: 137:88643

REFERENCE 7: 136:401651

REFERENCE 8: 135:376777

REFERENCE 9: 134:37033

REFERENCE 10: 133:359242

FILE 'HOME' ENTERED AT 10:33:19 ON 14 FEB 2003

Jiang, D.  
091756690 Page 1  
Seq. 1D 2

GenCore version 5.1.3  
Copyright (c) 1993 - 2003 Compugen Ltd.

OM protein - protein search, using SW model

Run on: February 13, 2003, 16:59:33 ; Search time 35 Seconds  
(without alignments)

148.479 Million cell updates/sec

Title: US-09-756-690a-2

Perfect score: 209

Sequence: 1 HGEETFTSDLSKQMEBEAVRLFTIWLNGGPSSGAPPPS 39

Scoring table: BLOSSOM62

Gapop 10.0 , Gapext 0.5

Searched: 908470 seqs, 13230620 residues

Total number of hits satisfying chosen parameters: 908470

Minimum DB seq length: 0

Maximum DB seq length: 200000000

Post-processing: Minimum Match 0%  
Maximum Match 10%  
Listing first 45 summaries

Database : A\_Geneset\_101002.\*

1: /SIDS2/gcddata/geneseq/geneseq-emb1/AA1980.DAT:\*

2: /SIDS2/gcddata/geneseq/geneseq-emb1/AA1981.DAT:\*

3: /SIDS2/gcddata/geneseq/geneseq-emb1/AA1982.DAT:\*

4: /SIDS2/gcddata/geneseq/geneseq-emb1/AA1983.DAT:\*

5: /SIDS2/gcddata/geneseq/geneseq-emb1/AA1984.DAT:\*

6: /SIDS2/gcddata/geneseq/geneseq-emb1/AA1985.DAT:\*

7: /SIDS2/gcddata/geneseq/geneseq-emb1/AA1986.DAT:\*

8: /SIDS2/gcddata/geneseq/geneseq-emb1/AA1987.DAT:\*

9: /SIDS2/gcddata/geneseq/geneseq-emb1/AA1988.DAT:\*

10: /SIDS2/gcddata/geneseq/geneseq-emb1/AA1989.DAT:\*

11: /SIDS2/gcddata/geneseq/geneseq-emb1/AA1990.DAT:\*

12: /SIDS2/gcddata/geneseq/geneseq-emb1/AA1991.DAT:\*

13: /SIDS2/gcddata/geneseq/geneseq-emb1/AA1992.DAT:\*

14: /SIDS2/gcddata/geneseq/geneseq-emb1/AA1993.DAT:\*

15: /SIDS2/gcddata/geneseq/geneseq-emb1/AA1994.DAT:\*

16: /SIDS2/gcddata/geneseq/geneseq-emb1/AA1995.DAT:\*

17: /SIDS2/gcddata/geneseq/geneseq-emb1/AA1996.DAT:\*

18: /SIDS2/gcddata/geneseq/geneseq-emb1/AA1997.DAT:\*

19: /SIDS2/gcddata/geneseq/geneseq-emb1/AA1998.DAT:\*

20: /SIDS2/gcddata/geneseq/geneseq-emb1/AA1999.DAT:\*

21: /SIDS2/gcddata/geneseq/geneseq-emb1/AA2000.DAT:\*

22: /SIDS2/gcddata/geneseq/geneseq-emb1/AA2001.DAT:\*

23: /SIDS2/gcddata/geneseq/geneseq-emb1/AA2002.DAT:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

## ALIGNMENTS

RESULT 1  
AAR80546 standard, peptide, 39 AA.  
ID AAR80546

AAR80546;

AC

DT

DE

Heloderma suspectum

Exendin-4; diabetes mellitus; hyperglycaemia; insulinotropic peptide.

## SUMMARIES

Result No.	Score	Query	Match	Length	DB ID	Description
1	209	100.0	39	16	AAR80546	Heloderma suspectu
2	209	100.0	39	16	AAR80546	Exendin-4, for use
3	203	100.0	39	16	AAR80546	Gila monster exend
4	209	100.0	39	20	AAY31502	Exendin-4 peptide
5	209	100.0	39	20	AAY31718	Amino acid sequenc
6	209	100.0	39	21	AAB11282	H. suspectum exend
7	203	100.0	39	21	AAB11305	Exendin agonist pe
8	209	100.0	39	21	AAB11306	Exendin agonist pe
9	209	100.0	39	21	AAB11307	Exendin agonist pe
10	209	100.0	39	21	AAB11308	Exendin agonist pe

PN US5424286-A.

PD 13-JUN-1995.

PF 24-MAY-1993;

PR 24-MAY-1993;

XX 93US-0066480.

XX 93US-0066480.

XX 93US-0066480.

Claim 6; Columns 13-14; 17pp; English.

-

Stimulating/inhibiting insulin release with exendin polypeptide(s) -

for treating diabetes mellitus and preventing hyperglycaemia.



RESULT 4  
 AAY31502  
 ID AAY31502 standard; peptide; 39 AA.  
 XX  
 AC AAY31502;  
 XX  
 DT 08-Nov-1999 (first entry)  
 XX  
 DE Exendin-4 peptide sequence.  
 XX  
 KW Exendin; agonist; GLP-1; glucagon-like peptide; toxic hypervolemia;  
 KW diureisis; renal plasma flow; glomerular filtration rate; pre-eclampsia;  
 KW eclampsia of pregnancy; cardiac contractility; renal failure; diuretic;  
 KW congestive heart failure; nephrotic syndrome; pulmonary edema; cirrhosis;  
 KW hypertension; urine flow.  
 XX  
 OS Synthetic.  
 OS Heloderma suspectum.  
 XX  
 PH Key  
 FT Modified-Site 39  
 FT /note= "C-terminal amide"  
 FT XX  
 PN WO940788-A1.  
 XX  
 PD 19-AUG-1999.  
 XX  
 PF 05-FEB-1999; 99WO-US02554.  
 XX  
 PR 13-FEB-1999; 98US-0075122.  
 XX  
 PA (AMYL-) AMYLIN PHARM INC.  
 XX  
 PI Beeley NRA, Prickett K, Vine W, Young AA;  
 XX  
 DR WPI; 1999-527332/44.  
 XX  
 PT Increasing urine flow by administering peptides or peptide agonists  
 XX  
 PS Claim 15: Page 7; 94pp; English.  
 XX  
 CC The invention relates to new methods of increasing urine flow that  
 CC comprises administering an exendin or exendin agonist, or a GLP-1  
 CC (glucagon-like peptide), or GLP-1 agonist. The new methods using an  
 CC exendin, exendin agonist, GLP-1 or GLP-1 agonist are useful for  
 CC increasing urine flow, decreasing potassium concentration in urine,  
 CC preventing or alleviating a disorder associated with toxic hypervolemia  
 CC (renal failure, congestive heart failure, nephritic syndrome, pulmonary  
 CC edema, cirrhosis, or hypertension). They can also be used for inducing  
 CC rapid diuresis, preparing an individual for surgical procedure,  
 CC increasing renal plasma flow and glomerular filtration rate, treating  
 CC pre-eclampsia or eclampsia of pregnancy, and increasing a condition/  
 CC disorder that can be alleviated by increasing cardiac contractility  
 CC (congestive heart failure, pulmonary edema, systemic edema, or renal  
 CC failure). Unlike prior art diuretics, the new methods increase urine  
 CC excretion and sodium excretion without increasing potassium loss, and are  
 CC fast acting. They have a prolonged duration of action, are inotropic,  
 CC have a low toxicity, and are easily administered intravenously. The  
 CC present sequence represents an exendin-4 peptide which can be used in  
 CC the methods of the invention.

Sequence 39 AA;  
 Query Match 100.0%; Score 209; DB 20; Length 39;  
 Best Local Similarity 100.0%; Pred. No. 2.3e-19;  
 Matches 39; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
 Qy 1 HGGCTFTSDLSKQMEBAVLFIEWLNGPSSGAPPPS 39  
 Db 1 HGGCTFTSDLSKQMEBAVLFIEWLNGPSSGAPPPS 39

RESULT 5  
 AAY03718 standard; peptide; 39 AA.  
 XX  
 AC AAY03718;  
 XX  
 DT 08-JUN-1999 (first entry)  
 XX  
 DE Amino acid sequence of exendin-4.  
 XX  
 KW Exendin; agonist; diabetes; disorder; plasma glucose; gastric;  
 KW diagnostic; gastro-intestinal; radiological; generic.  
 XX  
 OS Synthetic.  
 XX  
 PH Key  
 FT Modified-Site 39  
 FT /note= "C-terminal amide"  
 XX  
 PN WO9907404-A1.  
 XX  
 PD 18-FEB-1999.  
 XX  
 PR 06-AUG-1998; 98WO-US16387.  
 XX  
 PR 08-AUG-1997; 97US-0055404.  
 XX  
 PA (AMYL-) AMYLIN PHARM INC.  
 XX  
 PI Beeley NRA, Prickett KS;  
 XX  
 DR WPI; 1998-180403/15.  
 XX  
 PR New exendin agonists - useful in the treatment of Type I and II  
 diabetes.  
 XX  
 PS Disclosure; Fig 3; 70pp; English.  
 XX  
 CC The invention relates to exendin agonists which slow gastric emptying  
 CC and lower plasma glucose levels. The peptides are of the formula  
 CC Xaa1-Gly-Xaa2-Thr-Xaa4-Xaa5-Xaa6-Xaa7-Xaa8-Ser-Lys-Gln-Xaa9-Glu-Glu-  
 CC Val-Arg-Leu-Xaa10-Xaa11-Xaa12-Xaa13-Leu-Dys-Asn-Gly-Gly-Xaa14-Ser-Ser-  
 CC Gly-Ala-Xaa15-Xaa16-Xaa17-Xaa18-Z, wherein: Xaa1 is His, Arg or Tyr;  
 CC Xaa2 is Ser, Gly, Ala, or Thr; Xaa3 is Asp or Glu; Xaa4 is Phe, Tyr, or  
 CC naphthylalanine; Xaa5 is Thr or Ser; Xaa6 is Ser or Thr; Xaa7 is Asp or  
 CC Glu; Xaa8 is Leu, Ile, Val, Pentylglycine, or Met; Xaa9 is Leu, Ile,  
 CC pentylglycine, Val, or Met; Xaa10 is Phe, Tyr, or naphthylalanine; Xaa11  
 CC is Ile, Val, Leu, pentylglycine, tert-butylglycine, or Met; Xaa12 is Glu  
 CC or Asp; Xaa13 is Trp, Phe, Tyr, or naphthylalanine; Xaa14, Xaa15, Xaa16,  
 CC and Xaa17 are independently Pro, homopropylene, 3Hyp, 4Hyp, thiodipropylene,  
 CC Thr, or Tyr; and Z is -OH or -NH2 with the proviso that the sequence is  
 CC not the amino acid sequences shown in the present sequence and AAY03717.  
 CC The specification claims for a second peptide of the above formula where  
 CC Xaa1 is His, Arg, Tyr or 4-imidazolopropenyl. The exendin agonists are  
 CC used to treat Type I and II diabetes, disorders which would be benefited  
 CC by agents which lower plasma glucose levels, and disorders which would be  
 CC benefited by agents useful in delaying and/or slowing gastric emptying.  
 CC Delayed gastric emptying is a useful diagnostic aid in gastro-intestinal  
 CC radiological examinations. The present sequence represents the amino  
 CC acid sequence of exendin-4.  
 XX  
 SQ Sequence 39 AA;

Query Match 100.0%; Score 209; DB 20; Length 39;  
 Best Local Similarity 100.0%; Pred. No. 2.3e-19;  
 Matches 39; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
 Qy 1 HGGCTFTSDLSKQMEBAVLFIEWLNGPSSGAPPPS 39  
 Db 1 HGGCTFTSDLSKQMEBAVLFIEWLNGPSSGAPPPS 39

**RESULT 6**  
 AAB11282 ID AAB11282 standard; Peptide: 39 AA.  
 XX  
 AC AAB11282;  
 XX  
 DT 20-FEB-2001 (first entry)  
 DE H. suspectum exendin 4 peptide SEQ ID NO 2.  
 XX  
 KW Exendin; agonist; treatment; antidiabetic; insulin sensitivity; diabetes;  
 plasma glucose; gastric emptying; food intake.  
 XX  
 OS Heiderma suspectum.  
 XX  
 PN WO200041546-A2.  
 XX  
 PD 20-JUL-2000.  
 XX  
 PR 10-JAN-2000; 2000US-0116380.  
 XX  
 (AMYL-) AMYLIN PHARM INC.  
 XX  
 PI Young A, L'Italien JJ, Kolterman O;  
 XX  
 WPI; 2000-514584/46.  
 XX  
 New formulations comprising an exendin or exendin agonist peptide used  
 for increasing the sensitivity of a subject to insulin to treat  
 diabetes -  
 XX  
 PT  
 XX  
 PS Example 36; Figure 15; 281PP; English.  
 XX  
 This invention describes a novel formulation (I) comprising an exendin or  
 exendin agonist peptide, a buffer and an iso-osmolarity modifier which  
 has a pH of 3-7. The products of the invention have antidiabetic  
 activity. The exendin or exendin agonist is used to increase the  
 sensitivity of a subject to insulin to treat diabetes and disorders which  
 would benefit from agents which lower plasma glucose levels and disorders  
 which would benefit from agents that delay and/or slow gastric emptying  
 or reducing food intake.  
 XX  
 SQ Sequence 39 AA;  
 Query Match 100.0%; Score 209; DB 21; Length 39;  
 Best Local Similarity 100.0%; Pred. No. 2.3e-19;  
 Matches 39; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
 XX  
 PT  
 XX  
 Qy 1 HGEGETFTSDLSKQMEEEAVRLFIEWLKGPGSSGAPPS 39  
 Db 1 HGEGETFTSDLSKQMEEEAVRLFIEWLKGPGSSGAPPS 39  
 XX  
 PS Example 2; Figure 2; 281PP; English.  
 XX  
 This invention describes a novel formulation (I) comprising an exendin or  
 exendin agonist peptide, a buffer and an iso-osmolarity modifier which  
 has a pH of 3-7. The products of the invention have antidiabetic  
 activity. The exendin or exendin agonist is used to increase the  
 sensitivity of a subject to insulin to treat diabetes and disorders which  
 would benefit from agents which lower plasma glucose levels and disorders  
 which would benefit from agents that delay and/or slow gastric emptying  
 or reducing food intake.  
 XX  
 SQ Sequence 39 AA;  
 Query Match 100.0%; Score 209; DB 21; Length 39;  
 Best Local Similarity 100.0%; Pred. No. 2.3e-19;  
 Matches 39; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
 XX  
 PT  
 XX  
 Qy 1 HGEGETFTSDLSKQMEEEAVRLFIEWLKGPGSSGAPPS 39  
 Db 1 HGEGETFTSDLSKQMEEEAVRLFIEWLKGPGSSGAPPS 39  
 XX  
 PS Example 2; Figure 2; 281PP; English.  
 XX  
 This invention describes a novel formulation (I) comprising an exendin or  
 exendin agonist peptide, a buffer and an iso-osmolarity modifier which  
 has a pH of 3-7. The products of the invention have antidiabetic  
 activity. The exendin or exendin agonist is used to increase the  
 sensitivity of a subject to insulin to treat diabetes and disorders which  
 would benefit from agents which lower plasma glucose levels and disorders  
 which would benefit from agents that delay and/or slow gastric emptying  
 or reducing food intake.  
 XX  
 SQ Sequence 39 AA;  
 Query Match 100.0%; Score 209; DB 21; Length 39;  
 Best Local Similarity 100.0%; Pred. No. 2.3e-19;  
 Matches 39; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
 XX  
 PT  
 XX  
 Qy 1 HGEGETFTSDLSKQMEEEAVRLFIEWLKGPGSSGAPPS 39  
 Db 1 HGEGETFTSDLSKQMEEEAVRLFIEWLKGPGSSGAPPS 39  
 XX  
 PS Example 3; Figure 3; 281PP; English.  
 XX  
 This invention describes a novel formulation (I) comprising an exendin or  
 exendin agonist peptide, a buffer and an iso-osmolarity modifier which  
 has a pH of 3-7. The products of the invention have antidiabetic  
 activity. The exendin or exendin agonist is used to increase the  
 sensitivity of a subject to insulin to treat diabetes and disorders which  
 would benefit from agents which lower plasma glucose levels and disorders  
 which would benefit from agents that delay and/or slow gastric emptying  
 or reducing food intake.  
 XX  
 SQ Sequence 39 AA;  
 Query Match 100.0%; Score 209; DB 21; Length 39;  
 Best Local Similarity 100.0%; Pred. No. 2.3e-19;  
 Matches 39; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
 XX  
 PT  
 XX  
 Qy 1 HGEGETFTSDLSKQMEEEAVRLFIEWLKGPGSSGAPPS 39  
 Db 1 HGEGETFTSDLSKQMEEEAVRLFIEWLKGPGSSGAPPS 39  
 XX  
 PS Example 3; Figure 3; 281PP; English.

**RESULT 7**  
 AAB11305 ID AAB11305 standard; Peptide: 39 AA.  
 XX  
 AC AAB11305;  
 XX  
 DT 20-FEB-2001 (first entry)  
 XX  
 DE exendin agonist Peptide SEQ ID NO 31.  
 XX  
 KW Exendin; agonist; treatment; antidiabetic; insulin sensitivity; diabetes;  
 plasma glucose; gastric emptying; food intake.  
 OS Synthetic.  
 XX  
 PN WO200041546-A2.  
 XX  
 PD 20-JUL-2000.  
 XX  
 PR 10-JAN-2000; 2000US-0116380.  
 XX  
 (AMYL-) AMYLIN PHARM INC.  
 XX  
 PI Young A, L'Italien JJ, Kolterman O;  
 XX  
 WPI; 2000-514584/46.  
 XX  
 New formulations comprising an exendin or exendin agonist peptide used  
 for increasing the sensitivity of a subject to insulin to treat  
 diabetes -  
 XX  
 PT  
 XX  
 PS Example 37; Figure 15; 281PP; English.

XX This invention describes a novel formulation (I) comprising an exendin or exendin agonist peptide, a buffer and an iso-osmolarity modifier which has a pH of 3-7. The products of the invention have antidiabetic activity. The exendin or exendin agonist is used to increase the sensitivity of a subject to insulin to treat diabetes and disorders which would benefit from agents which lower plasma glucose levels and disorders which would benefit from agents that delay and/or slow gastric emptying or reducing food intake.

XX Sequence 39 AA;

Query Match 100.0%; Score 209; DB 21; Length 39;

Best Local Similarity 100.0%; Pred. No. 2.3e-19; Matches 39; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

PS 1 HGEGLFTSDLSKQMEBAAVLFIENLKGPGSSGAPPS 39

RESULT 9

AB11307 ID AAB11307 standard; Peptide; 39 AA.

XX

AC AAB11307;

XX

DT 20-FEB-2001 (first entry)

XX

DB exendin agonist peptide SEQ ID NO 33.

XX

RW Exendin; agonist; treatment; antidiabetic; insulin sensitivity; diabetes; plasma glucose; gastric emptying; food intake.

XX

OS Synthetic.

XX

PS WO200041546-A2.

XX

PD 20-JUL-2000.

XX

PP 10-JAN-2000; 2000US-0116380.

XX

PR 14-JAN-1999; 99US-0116380.

XX

PA (AMYL-) AMYLIN PHARM INC.

XX

PI Young A, L'Italien JJ, Kolterman O;

XX

DR WPI; 2000-514584/46.

XX

PS Example 39; Figure 15; 281PP; English.

XX

CC This invention describes a novel formulation (I) comprising an exendin or exendin agonist peptide, a buffer and an iso-osmolarity modifier which has a pH of 3-7. The products of the invention have antidiabetic activity. The exendin or exendin agonist is used to increase the sensitivity of a subject to insulin to treat diabetes and disorders which would benefit from agents which lower plasma glucose levels and disorders which would benefit from agents that delay and/or slow gastric emptying or reducing food intake.

XX Sequence 39 AA;

Query Match 100.0%; Score 209; DB 21; Length 39;

Best Local Similarity 100.0%; Pred. No. 2.3e-19; Matches 39; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

PS 1 HGEGLFTSDLSKQMEBAAVLFIENLKGPGSSGAPPS 39

XX

DB 1 HGEGLFTSDLSKQMEBAAVLFIENLKGPGSSGAPPS 39

XX

DE Extendin-4 peptide #7.

XX

RW Extendin; agonist; diabetes; obesity; eating disorder;

XX dyslipidaemia; insulin-resistance syndrome; food intake.

XX OS Heloderma sp.

Db 1 HGEGLFTSDLSKQMEBAAVLFIENLKGPGSSGAPPS 39

RESULT 10  
AAB11308 ID AAB11308 standard; Peptide; 39 AA.  
XX  
AC AAB11308;  
XX  
DT 20-FEB-2001 (first entry)

DB exendin agonist peptide SEQ ID NO 34.

XX

RW Exendin; agonist; treatment; antidiabetic; insulin sensitivity; diabetes; plasma glucose; gastric emptying; food intake.

XX

OS Synthetic.

XX

PS WO200041546-A2.

XX

PD 20-JUL-2000.

XX

PP 10-JAN-2000; 2000US-0116380.

XX

PR 14-JAN-1999; 99US-0116380.

XX

PA (AMYL-) AMYLIN PHARM INC.

XX

PI Young A, L'Italien JJ, Kolterman O;

XX

DR WPI; 2000-514584/46.

XX

PS Example 39; Figure 15; 281PP; English.

XX

CC This invention describes a novel formulation (I) comprising an exendin or exendin agonist peptide, a buffer and an iso-osmolarity modifier which has a pH of 3-7. The products of the invention have antidiabetic activity. The exendin or exendin agonist is used to increase the sensitivity of a subject to insulin to treat diabetes and disorders which would benefit from agents which lower plasma glucose levels and disorders which would benefit from agents that delay and/or slow gastric emptying or reducing food intake.

XX Sequence 39 AA;

Query Match 100.0%; Score 209; DB 21; Length 39;

Best Local Similarity 100.0%; Pred. No. 2.3e-19; Matches 39; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

PS 1 HGEGLFTSDLSKQMEBAAVLFIENLKGPGSSGAPPS 39

XX

DB 1 HGEGLFTSDLSKQMEBAAVLFIENLKGPGSSGAPPS 39

XX

DE Extendin-4 peptide #7.

XX

RW Extendin; agonist; diabetes; obesity; eating disorder;

XX dyslipidaemia; insulin-resistance syndrome; food intake.

XX OS Heloderma sp.

Db 1 HGEGLFTSDLSKQMEBAAVLFIENLKGPGSSGAPPS 39

RESULT 11  
AAB52840 ID AAB52840 standard; Peptide; 39 AA.  
XX  
AC AAB52840;  
XX  
DT 28-FEB-2001 (first entry)

DB Extendin-4 peptide #7.

XX

RW Extendin; agonist; diabetes; obesity; eating disorder;

XX dyslipidaemia; insulin-resistance syndrome; food intake.

XX OS Heloderma sp.

Db 1 HGEGLFTSDLSKQMEBAAVLFIENLKGPGSSGAPPS 39

PN WO200066629-A1.  
 XX 09-NOV-2000.  
 XX 28-APR-2000; 2000WO-US11814.  
 PF PR 30-APR-1999; 99US-0132018.  
 XX PA (ANYL-) ANYLIN PHARM INC.  
 XX Young A, Prickett K;  
 XX DR WPI; 2000-672834/65.  
 PT Modified exendin or an exendin agonist linked to one or more polyethylene glycol (PEG) polymers, modulate plasma glucose levels, useful for treating disorders such as diabetes and obesity -

Example 4; Page 71; 119PP; English.

XX The present invention relates to exendins and their agonists which have been modified with molecular weight increasing agents such as polyethylene glycol (PEG). These can be used in the treatment of diabetes, obesity, impaired glucose tolerance, postprandial dumping syndrome, postprandial hyperglycaemia, eating disorders, insulin resistance syndrome, dyslipidaemia and to suppress glucagon secretion.

Sequence 39 AA;

Query Match 100.0%; Score 209; DB 21; Length 39;  
 Best Local Similarity 100.0%; Pred. No. 2.3e-19;  
 Matches 39; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
 Qy 1 HEGEGFTSDLSKQMEBEAVRLFIEWLNGGPGSSGAPPPS 39  
 Db 1 HGGTFTSDLSKQMEBEAVRLFIEWLNGGPGSSGAPPPS 39

RESULT 13

AAB52857 standard; Peptide; 39 AA.  
 ID AAB52857  
 XX AC AA52857;  
 XX DT 28-FEB-2001 (first entry)  
 XX DB Extendin-4 peptide #24.  
 XX KW Extendin; agonist; diabetes; obesity; eating disorder;  
 XX KW dyslipidaemia; insulin-resistance syndrome; food intake.  
 XX OS Heloderma sp.  
 PN WO200066629-A1.  
 PD 09-NOV-2000.  
 XX PF 28-APR-2000; 2000WO-US11814.  
 XX PR 30-APR-1999; 99US-0132018.  
 XX PA (ANYL-) ANYLIN PHARM INC.  
 XX PI Young A, Prickett K;  
 XX DR WPI; 2000-672834/65.

XX Extendin-4 peptide #8.

XX Extendin; agonist; diabetes; obesity; eating disorder;  
 KW dyslipidaemia; insulin-resistance syndrome; food intake.  
 XX OS Heloderma sp.  
 XX PN WO200066629-A1.  
 XX 09-NOV-2000.  
 PF XX 28-APR-2000; 2000WO-US11814.  
 XX PR 30-APR-1999; 99US-0132018.  
 XX PA (ANYL-) ANYLIN PHARM INC.

XX Young A, Prickett K;  
 XX DR WPI; 2000-672834/65.  
 PT Modified exendin or an exendin agonist linked to one or more polyethylene glycol (PEG) polymers, modulate plasma glucose levels, useful for treating disorders such as diabetes and obesity -

Sequence 39 AA;

Query Match 100.0%; Score 209; DB 21; Length 39;  
 Best Local Similarity 100.0%; Pred. No. 2.3e-19;  
 Matches 39; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
 Qy 1 HEGEGFTSDLSKQMEBEAVRLFIEWLNGGPGSSGAPPPS 39  
 Db 1 HGGTFTSDLSKQMEBEAVRLFIEWLNGGPGSSGAPPPS 39

RESULT 14

AAB52858

CC The present invention relates to exendins and their agonists which have been modified with molecular weight increasing agents such as polyethylene glycol (PEG). These can be used in the treatment of diabetes, obesity, impaired glucose tolerance, postprandial dumping syndrome, postprandial hyperglycaemia, eating disorders, insulin resistance syndrome, dyslipidaemia and to suppress glucagon secretion.

Sequence 39 AA;

Query Match 100.0%; Score 209; DB 21; Length 39;  
 Best Local Similarity 100.0%; Pred. No. 2.3e-19;  
 Matches 39; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
 Qy 1 HEGEGFTSDLSKQMEBEAVRLFIEWLNGGPGSSGAPPPS 39  
 Db 1 HGGTFTSDLSKQMEBEAVRLFIEWLNGGPGSSGAPPPS 39

RESULT 13

AAB52857 standard; Peptide; 39 AA.  
 ID AAB52857  
 XX AC AA52857;  
 XX DT 28-FEB-2001 (first entry)  
 XX DB Extendin-4 peptide #24.  
 XX KW Extendin; agonist; diabetes; obesity; eating disorder;  
 XX KW dyslipidaemia; insulin-resistance syndrome; food intake.  
 XX OS Heloderma sp.  
 PN WO200066629-A1.  
 PD 09-NOV-2000.  
 XX PF 28-APR-2000; 2000WO-US11814.  
 XX PR 30-APR-1999; 99US-0132018.  
 XX PA (ANYL-) ANYLIN PHARM INC.  
 XX PI Young A, Prickett K;  
 XX DR WPI; 2000-672834/65.

XX Extendin-4 peptide #8.

XX Extendin; agonist; diabetes; obesity; eating disorder;  
 KW dyslipidaemia; insulin-resistance syndrome; food intake.  
 XX OS Heloderma sp.  
 XX PN WO200066629-A1.  
 XX 09-NOV-2000.  
 PF XX 28-APR-2000; 2000WO-US11814.  
 XX PR 30-APR-1999; 99US-0132018.  
 XX PA (ANYL-) ANYLIN PHARM INC.

XX Young A, Prickett K;  
 XX DR WPI; 2000-672834/65.  
 PT Modified exendin or an exendin agonist linked to one or more polyethylene glycol (PEG) polymers, modulate plasma glucose levels, useful for treating disorders such as diabetes and obesity -

Sequence 39 AA;

Query Match 100.0%; Score 209; DB 21; Length 39;  
 Best Local Similarity 100.0%; Pred. No. 2.3e-19;  
 Matches 39; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
 Qy 1 HEGEGFTSDLSKQMEBEAVRLFIEWLNGGPGSSGAPPPS 39  
 Db 1 HGGTFTSDLSKQMEBEAVRLFIEWLNGGPGSSGAPPPS 39

RESULT 14

AAB52858

ID AAB52858 standard; Peptide; 39 AA.  
 XX  
 AC AAB52858;  
 XX DT 28-FEB-2001 (first entry)  
 XX DE Extendin-4 peptide #25.  
 XX KW Extendin; agonist; diabetes; eating disorder;  
 KW dyslipidaemia; insulin-resistance syndrome; food intake.  
 XX OS Heloderma sp.  
 XX PN WO200066629-A1.  
 XX PD 09-NOV-2000.  
 XX PT 28-APR-2000; 2000WO-US11814.  
 XX PR 30-APR-1999; 99US-0132018.  
 XX PA (AMYL-) AMYLIN PHARM INC.  
 PI Young A, Prickett K;  
 XX DR 2000-672834/65.  
 XX  
 The present invention relates to extendins and their agonists which have  
 been modified with molecular weight increasing agents such as  
 polyethylene glycol (PEG). These can be used in the treatment of  
 diabetes, obesity, impaired glucose tolerance, postprandial dumping  
 syndrome, postprandial hyperglycaemia, eating disorders, insulin  
 resistance syndrome, dyslipidaemia and to suppress glucagon secretion.  
 XX  
 Example 2; Fig 2; 119pp; English.  
 XX  
 The present invention relates to extendins and their agonists which have  
 been modified with molecular weight increasing agents such as  
 polyethylene glycol (PEG). These can be used in the treatment of  
 diabetes, obesity, impaired glucose tolerance, postprandial dumping  
 syndrome, postprandial hyperglycaemia, eating disorders, insulin  
 resistance syndrome, dyslipidaemia and to suppress glucagon secretion.  
 XX  
 Example 2; Fig 2; 119pp; English.  
 XX  
 The present invention relates to extendins and their agonists which have  
 been modified with molecular weight increasing agents such as  
 polyethylene glycol (PEG). These can be used in the treatment of  
 diabetes, obesity, impaired glucose tolerance, postprandial dumping  
 syndrome, postprandial hyperglycaemia, eating disorders, insulin  
 resistance syndrome, dyslipidaemia and to suppress glucagon secretion.  
 XX  
 Sequence 39 AA;  
 Query Match 100.0%; Score 209; DB 21; Length 39;  
 Best Local Similarity 100.0%; Pred. No. 2.3e-19;  
 Matches 39; Conservative 0; Mismatches 0;  
 Indels 0; Gaps 0;  
 XX  
 Qy 1 HGEGTFTSDLSKQMBEAVRLFIEWLNGGPPSSGAPPPS 39  
 Db 1 HGEGTFTSDLSKQMBEAVRLFIEWLNGGSSGAPPPS 39  
 XX  
 Search completed: February 13, 2003, 17:10:47  
 Job time : 36 secs

ID AAB52872 standard; Peptide; 39 AA.  
 XX  
 AC AAB52872;  
 XX DT 28-FEB-2001 (first entry)  
 XX DE Gila monster extendin-4 protein.  
 XX KW Extendin; agonist; diabetes; obesity; eating disorder;  
 KW dyslipidaemia; insulin-resistance syndrome; food intake.  
 XX OS Heloderma suspectum.  
 XX PN WO200066629-A1.  
 XX PD 09-NOV-2000.  
 XX PF 28-APR-2000; 2000WO-US11814.  
 XX PR 30-APR-1999; 99US-0132018.

RESULT 15  
 AAB52872  
 ID AAB52872 standard; Peptide; 39 AA.  
 XX  
 AAB52872;  
 AC  
 XX  
 DE Gila monster extendin-4 protein.  
 XX KW Extendin; agonist; diabetes; obesity; eating disorder;  
 KW dyslipidaemia; insulin-resistance syndrome; food intake.  
 XX OS Heloderma suspectum.  
 XX PN WO200066629-A1.  
 XX PD 09-NOV-2000.  
 XX PF 28-APR-2000; 2000WO-US11814.  
 XX PR 30-APR-1999; 99US-0132018.



Db 1 HGGTFTSDLSKQMEEAIRLPIEWLKGPGSSGAPPS 39

RESULT 2  
US-09-302-596-9

; Sequence 9, Application US/09302596

; GENERAL INFORMATION:

; APPLICANT: Coolidge, Thomas R.

; ATTORNEY: Bhatia, Mario R.W.

; TITLE OF INVENTION: Metabolic Intervention with GLP-1 to Improve the Function of

; Title of Invention: Metabolic Intervention with GLP-1 to Improve the Function of

; CURRENT APPLICATION NUMBER: US/09/302,596

; CURRENT FILING DATE: 1999-04-30

; PRIOR APPLICATION NUMBER: 60/103,498

; PRIOR FILING DATE: 1998-10-08

; NUMBER OF SEQ ID NOS: 13

; SOFTWARE: PatentIn Ver. 2.0

; SEQ ID NO: 9

; LENGTH: 39

; TYPE: PRT

; ORGANISM: Gila Monster venom

US-09-302-596-9

Query Match 100.0%; Score 209; DB 4; Length 39;

Best Local Similarity 100.0%; Pred. No. 2.6e-20;

Matches 39; Conservative 0; Mismatches 0;

Indels 0; Gaps 0;

Qy 1 HGGTFTSDLSKQMEEAIRLPIEWLKGPGSSGAPPS 39

Db 1 HGGTFTSDLSKQMEEAIRLPIEWLKGPGSSGAPPS 39

RESULT 3

US-09-623-618B-12

; Sequence 12, Application US/09623618B

; GENERAL INFORMATION:

; APPLICANT: Bridon, Dominique P.

; ATTORNEY: L'Archeveque, Benoit

; APPLICANT: Ezrin, Alan M.

; APPLICANT: Holmes, Darren L.

; APPLICANT: Leblanc, Anouk

; APPLICANT: St. Pierre, Serge

; TITLE OF INVENTION: LONG LASTING INSULINOTROPIC PEPTIDES

; FILE REFERENCE: 501082001620

; CURRENT APPLICATION NUMBER: US/09/623,618B

; CURRENT FILING DATE: 2000-09-05

; PRIOR APPLICATION NUMBER: PCT/US00/13563

; PRIOR FILING DATE: 2000-05-17

; PRIOR APPLICATION NUMBER: 60/159,783

; PRIOR FILING DATE: 1999-10-15

; PRIOR APPLICATION NUMBER: 60/134,406

; PRIOR FILING DATE: 1999-05-17

; NUMBER OF SEQ ID NOS: 35

; SOFTWARE: FastSEQ for Windows Version 4.0

; SEQ ID NO: 12

; LENGTH: 39

; TYPE: PRT

; ORGANISM: Artificial Sequence

; FEATURE:

; OTHER INFORMATION: Description of Artificial Sequence: Synthetic

US-09-623-618B-12

Query Match 100.0%; Score 209; DB 4; Length 39;

Best Local Similarity 100.0%; Pred. No. 2.6e-20;

Matches 39; Conservative 0; Mismatches 0;

Indels 0; Gaps 0;

Qy 1 HGGTFTSDLSKQMEEAIRLPIEWLKGPGSSGAPPS 39

Db 1 HGGTFTSDLSKQMEEAIRLPIEWLKGPGSSGAPPS 39

RESULT 5

US-09-303-016-9

; Sequence 9, Application US/09303016

; Patent No. 6429,97

; GENERAL INFORMATION:

; APPLICANT: Coolidge, Thomas R.

; ATTORNEY: Bhiers, Mario R.W.

; TITLE OF INVENTION: Metabolic Intervention with GLP-1 or its Biologically

; Active Analogues to Improve the Function of the

; TITLE OF INVENTION: Ischemic and Reperfused Brain

; FILE REFERENCE: P03660US2

; CURRENT APPLICATION NUMBER: US/09/303,016

; CURRENT FILING DATE: 1999-04-30

; PRIOR APPLICATION NUMBER: 60/103,498

; PRIOR FILING DATE: 1998-10-08

; NUMBER OF SEQ ID NOS: 13

; SOFTWARE: PatentIn Ver. 2.0

; SEQ ID NO: 9

; LENGTH: 39

; TYPE: PRT

; ORGANISM: Heloderma suspectum

US-09-303-016-9

Query Match 100.0%; Score 209; DB 4; Length 39;

Best Local Similarity 100.0%; Pred. No. 2.6e-20;

Matches 39; Conservative 0; Mismatches 0;

Indels 0; Gaps 0;

Qy 1 HGGTFTSDLSKQMEEAIRLPIEWLKGPGSSGAPPS 39

Db 1 HGGTFTSDLSKQMEEAIRLPIEWLKGPGSSGAPPS 39

RESULT 4

US-09-333-415-9

; Sequence 9, Application US/09333415

; Patent No. 634180

; GENERAL INFORMATION:

; APPLICANT: Holst, Jens J.

; ATTORNEY: Vilbøll, Tina

; TITLE OF INVENTION: GLP-1 as a Diagnostic Test to Determine Beta-Cell

; Function and the Presence of the Condition of IGT and

; Type-II Diabetes

; FILE REFERENCE: P03987US0

; CURRENT APPLICATION NUMBER: US/09/333,415

; CURRENT FILING DATE: 1999-06-15

; NUMBER OF SEQ ID NOS: 13

; SOFTWARE: PatentIn Ver. 2.0

; SEQ ID NO: 9

; LENGTH: 39

; TYPE: PRT

; ORGANISM: Heloderma suspectum

US-09-333-415-9

Query Match 100.0%; Score 209; DB 4; Length 39;

Best Local Similarity 100.0%; Pred. No. 2.6e-20;

Matches 39; Conservative 0; Mismatches 0;

Indels 0; Gaps 0;

Qy 1 HGGTFTSDLSKQMEEAIRLPIEWLKGPGSSGAPPS 39

Db 1 HGGTFTSDLSKQMEEAIRLPIEWLKGPGSSGAPPS 39

RESULT 5

US-09-303-016-9

; Sequence 9, Application US/09303016

; Patent No. 6429,97

; GENERAL INFORMATION:

; APPLICANT: Coolidge, Thomas R.

; ATTORNEY: Bhiers, Mario R.W.

; TITLE OF INVENTION: Metabolic Intervention with GLP-1 or its Biologically

; Active Analogues to Improve the Function of the

; TITLE OF INVENTION: Ischemic and Reperfused Brain

; FILE REFERENCE: P03660US2

; CURRENT APPLICATION NUMBER: US/09/303,016

; CURRENT FILING DATE: 1999-04-30

; PRIOR APPLICATION NUMBER: 60/103,498

; PRIOR FILING DATE: 1998-10-08

; NUMBER OF SEQ ID NOS: 13

; SOFTWARE: PatentIn Ver. 2.0

; SEQ ID NO: 9

; LENGTH: 39

; TYPE: PRT

; ORGANISM: Heloderma suspectum

US-09-303-016-9

Query Match 100.0%; Score 209; DB 4; Length 39;

Best Local Similarity 100.0%; Pred. No. 2.6e-20;

Matches 39; Conservative 0; Mismatches 0;

Indels 0; Gaps 0;

Qy 1 HGGTFTSDLSKQMEEAIRLPIEWLKGPGSSGAPPS 39

Db 1 HGGTFTSDLSKQMEEAIRLPIEWLKGPGSSGAPPS 39

RESULT 6

US-09-623-618B-18

; Sequence 18, Application US/09623618B

; Patent No. 6329336

; GENERAL INFORMATION:

; APPLICANT: Bridon, Dominique P.

; ATTORNEY: L'Archeveque, Benoit

; TITLE OF INVENTION: Peptide

; FILE REFERENCE: Ezrin, Alan M.

; CURRENT APPLICATION NUMBER: US/09/623,618B

; CURRENT FILING DATE: 2000-09-05

; PRIOR APPLICATION NUMBER: PCT/US00/13563

; PRIOR FILING DATE: 2000-05-17

; NUMBER OF SEQ ID NOS: 35

; SOFTWARE: FastSEQ for Windows Version 4.0

; SEQ ID NO: 12

; LENGTH: 39

; TYPE: PRT

; ORGANISM: Artificial Sequence

US-09-623-618B-18

; Sequence 18, Application US/09623618B

; Patent No. 6329336

; GENERAL INFORMATION:

; APPLICANT: Bridon, Dominique P.

; ATTORNEY: L'Archeveque, Benoit

; TITLE OF INVENTION: Peptide

; FILE REFERENCE: Ezrin, Alan M.

APPLICANT: Holmes, Darren L.  
APPLICANT: Leblanc, Anouk  
APPLICANT: St. Pierre, Serge  
TITLE OF INVENTION: LONG LASTING INSULINOTROPIC PEPTIDES  
FILE REFERENCE: 500862001620  
CURRENT APPLICATION NUMBER: US/09/623,616B  
CURRENT FILING DATE: 2000-09-05  
PRIOR APPLICATION NUMBER: PCT/US00/15563  
PRIOR FILING DATE: 2000-05-17  
PRIOR APPLICATION NUMBER: 60/159,783  
PRIOR FILING DATE: 1999-10-15  
PRIOR APPLICATION NUMBER: 60/134,406  
PRIOR FILING DATE: 1999-05-17  
NUMBER OF SEQ ID NOS: 35  
SOFTWARE: FastSEQ for Windows Version 4.0  
SEQ ID NO: 18  
LENGTH: 40  
TYPE: PRT  
ORGANISM: Artificial Sequence  
FEATURE: OTHER INFORMATION: Description of Artificial Sequence : Synthetic  
OTHER INFORMATION: Peptide  
US-6-23,619-10

```

5-323-010-1a

Query Match          100.0%; Score 209; DB 4; Length 40;
Best Local Similarity 100.0%; Pred. No. 2, 7e-20;
Matches 39; Conservative 0; Mismatches 0; Indels 0;
1 HGBOTFTPSDSLSCOMEBAEAVRLPFBWLNKGQPSGAPPSS 39
1 HGBOTFTPSDSLSCOMEBAEAVRLPFBWLNKGQPSGAPPSS 39

```

RESULT 7  
IS-09-623-618B-31  
Sequence 31, Application US/0962361B  
Patent No. 6323336  
GENERAL INFORMATION:  
APPLICANT: L'Archeveque, Dominique P.  
APPLICANT: L'Archeveque, Benoit

APPLICANT: Brin, Alan M.  
APPLICANT: Holmes, Darren L.  
APPLICANT: Lablanc, Anouk  
APPLICANT: St. Pierre, Serge  
TITLE OF INVENTION: LONG LASTING INSULINOTROPIC PEPTIDES  
FILE REFERENCE: 500862001620  
CURRENT APPLICATION NUMBER: US/09/623,618B  
CURRENT FILING DATE: 2000-09-05  
PRIOR APPLICATION NUMBER: PCT/US/00/13563

PRIOR FILING DATE: 2000-05-17  
PRIOR APPLICATION NUMBER: 60/159,783  
PRIOR FILING DATE: 1999-10-15  
PRIOR APPLICATION NUMBER: 60/134,406  
PRIOR FILING DATE: 1999-05-17  
NUMBER OF SEQ ID NOS: 35  
SOFTWARE: FastSEQ for Windows Version 4.0  
SEQ ID NO: 31  
LENGTH: 40

TYPE: PRT  
 ORGANISM: Artificial Sequence  
 FEATURE: OTHER INFORMATION: Description of Artificial Sequence: Synthesis  
 OTHER INFORMATION: Peptide  
 NAME/KEY: MOD\_RES  
 LOCATION: 40  
 OTHER INFORMATION: Xaa represents Lys(E-NPA) - NH2-5TFA and where  
 IS-09-623-618B-31

Query Match 100.0% Score 209, DB 4;  
 Best Local Similarity 100.0% Pred. No. 2.7e-20  
 Matches 39; Conservative 0; Mismatches 0

1. HIGEGETTSDSLKOMEAVRLITEMLKGNGPSSGAPPPS 39

Db 1 FEGGFTSDLSKQMEEEAVRLFTEWLKNGGSSGAPPS 39

RESULT 8  
US-09-623-618B-32  
Sequence 32, Application US/09623618B  
Patent No. 6129336

GENERAL INFORMATION:

APPLICANT: Bridon, Dominique P.  
APPLICANT: L'Archeveque, Benoit  
APPLICANT: Ezrin, Alan M.  
APPLICANT: Holmes, Darren L.  
APPLICANT: Leblanc, Anouk  
APPLICANT: St. Pierre, Serge

TITLE OF INVENTION: LONG LASTING INSULINOTROPIC PE  
FILE REFERENCE: 50082001620

CURRENT APPLICATION NUMBER: US/09/623-618B  
CURRENT FILING DATE: 2000-09-05

PRIOR APPLICATION NUMBER: PCT/US00/113563  
PRIOR FILING DATE: 2000-05-17

PRIOR APPLICATION NUMBER: 60/159,783  
PRIOR FILING DATE: 2000-10-15

PRIOR FILING DATE: 12/27/1993-10-15  
PRIOR APPLICATION NUMBER: 60/1134,406  
PRIOR FILING DATE: 1999-05-17  
NUMBER OF SEQ ID NOS: 35  
SOFTWARE: FastSEQ for Windows Version 4.0  
SEQ ID NO: 32  
LENGTH: 40  
TYPE: PRT  
ORGANISM: Artificial Sequence

FEATURE: Description of Artificial Sequences  
OTHER INFORMATION: Description of Artificial Sequences  
OTHER INFORMATION: Peptide  
NAME/KEY: MOD\_RES  
LOCATION: 40  
OTHER INFORMATION: Xaa represents Lys(E)-AEEA-AEEA  
US-09-623-5188-32

```

Best Local Similarity 100.0%; Pred. No. 2.7e-20;
Matches 35; Conservative 0; Mismatches 0;
Indels 0; Gaps 0;

Qy 1 HGBGTFTSDLQMEEARVLFIRLFWLKNQGPSSGAPPS 39
Db 1 HGBGTFTSDLQMEEARVLFIRLFWLKNQGPSSGAPPS 39

```

US-066-480-1  
; Sequence 1, Application US/08066480  
; Patent No. 5424386  
; GENERAL INFORMATION:  
; APPLICANT: Eng, John  
; TITLE OF INVENTION: Pharmaceutical Compositions And Use of  
; Exendin-3 and Exendin-4 for Treatment of Diabetes Mellitus  
; NUMBER OF SEQUENCES: 7  
; CORRESPONDENCE ADDRESS:

1 ADDRESS: Allegretti & Witcoff, Ltd.  
1 STREET: 10 S. Wacker Drive  
1 CITY: Chicago  
1 STATE: Illinois  
1 COUNTRY: USA  
1 ZIP: 60606  
1 COMPUTER READABLE FORM:  
1 MEDIUM TYPE: Floppy disk  
1 COMPUTER: IBM PC compatible  
1 OPERATING SYSTEM: PC-DOS/MS-DOS  
1 SOFTWARE: PatentIn Release #1.0, Version #1.25  
1 CURRENT APPLICATION DATA:  
1 APPLICATION NUMBER: US/08/066,480  
1 FILING DATE: 24-MAR-1993  
1 CLASSIFICATION: 514

ATTORNEY/AGENT INFORMATION:  
; NAME: McDonnell, John J.  
; REGISTRATION NUMBER: 26,949  
; REFERENCE/DOCKET NUMBER: 93,084  
; TELECOMMUNICATION INFORMATION:  
; TELEPHONE: 312-715-1000  
; TELEFAX: 312-715-1234  
; INFORMATION FOR SEQ ID NO: 1:  
; SEQUENCE CHARACTERISTICS:  
; LENGTH: 39 amino acids  
; TYPE: amino acid  
; STRANDEDNESS: single  
; TOPOLOGY: linear  
; MOLECULE TYPE: peptide  
; FEATURE:  
; NAME/KEY: Peptide  
; LOCATION: 1..39  
; OTHER INFORMATION: /label= Exendin-3

US-08-066-180-1

Query Match 95.7%; Score 200; DB 1; Length 39;  
Best Local Similarity 94.9%; Pred. No. 3..76-19;  
Matches 37; Conservative 1; Mismatches 1; Indels 0; Gaps 0;

Qy 1 HGGTFTSDLSKQMEAVRLFIEWLNGKGPSSGAPPPS 39  
Db 1 HSDGFTFTSDLSKQMEAVRLFIEWLNGKGPSSGAPPPS 39

RESULT 10  
US-09-302-596-7  
; Sequence 7, Application US/09302596  
; PATENT NO. 6284725  
; GENERAL INFORMATION:  
; APPLICANT: Coolidge, Thomas R.  
; APPLICANT: Ehlers, Mario R.  
; TITLE OF INVENTION: Metabolic Intervention with GLP-1 to Improve the Function of  
; TITLE OF INVENTION: Ischemic and Reperfused Tissue  
; FILE REFERENCE: P3660US1  
; CURRENT APPLICATION NUMBER: US/09/302,596  
; CURRENT FILING DATE: 1999-06-30  
; PRIOR APPLICATION NUMBER: 60/103,498  
; PRIOR FILING DATE: 1998-10-08  
; SOFTWARE: PatentIn Ver. 2.0  
; SEQ ID NO 7  
; LENGTH: 39  
; TYPE: PAT  
; ORGANISM: Gila Monster venom

US-09-302-536-7

Query Match 95.7%; Score 200; DB 4; Length 39;  
Best Local Similarity 94.9%; Pred. No. 3..76-19;  
Matches 37; Conservative 1; Mismatches 1; Indels 0; Gaps 0;

Qy 1 HGGTFTSDLSKQMEAVRLFIEWLNGKGPSSGAPPPS 39  
Db 1 HSDGFTFTSDLSKQMEAVRLFIEWLNGKGPSSGAPPPS 39

RESULT 11  
US-09-623-618B-11  
; Sequence 11, Application US/09623618B  
; PATENT NO. 6329336  
; GENERAL INFORMATION:  
; APPLICANT: Bridon, Dominique P.  
; APPLICANT: L'Archeveque, Benoit  
; APPLICANT: Ezrin, Alan M.  
; APPLICANT: Holmes, Darren L.  
; APPLICANT: Leblanc, Anouk  
; APPLICANT: St. Pierre, Serge  
; TITLE OF INVENTION: LONG LASTING INSULINOTROPIC PEPTIDES  
; FILE REFERENCE: 50086201620

CURRENT APPLICATION NUMBER: US/09/623,618B  
; CURRENT FILING DATE: 2000-09-05  
; PRIOR APPLICATION NUMBER: PCT/US00/13563  
; PRIOR FILING DATE: 2000-05-17  
; PRIOR APPLICATION NUMBER: 60/159,783  
; PRIOR FILING DATE: 1999-10-15  
; PRIOR APPLICATION NUMBER: 60/134,406  
; PRIOR FILING DATE: 1999-05-17  
; NUMBER OF SEQ ID NOS: 35  
; SEQ ID NO 11  
; SOFTWARE: FastSEQ for Windows Version 4.0  
; LENGTH: 39  
; TYPE: PRF  
; ORGANISM: Artificial Sequence  
; FEATURE:  
; OTHER INFORMATION: Description of Artificial Sequence: Synthetic  
; OTHER INFORMATION: Peptide  
; OTHER INFORMATION: Peptide  
US-09-623-618B-11

Query Match 95.7%; Score 200; DB 4; Length 39;  
Best Local Similarity 94.9%; Pred. No. 3..76-19;  
Matches 37; Conservative 1; Mismatches 1; Indels 0; Gaps 0;

Qy 1 HGGTFTSDLSKQMEAVRLFIEWLNGKGPSSGAPPPS 39  
Db 1 HSDGFTFTSDLSKQMEAVRLFIEWLNGKGPSSGAPPPS 39

RESULT 12  
US-09-333-415-7  
; Sequence 7, Application US/09333415  
; PATENT NO. 634180  
; GENERAL INFORMATION:  
; APPLICANT: Valsboll, Tina  
; APPLICANT: Holst, Jens J.  
; TITLE OF INVENTION: GLP-1 as a Diagnostic Test to Determine Beta-Cell  
; Function and the Presence of the Condition of IGT and  
; Title: Type-II Diabetes  
; FILE REFERENCE: P03987US0  
; CURRENT APPLICATION NUMBER: US/09/333,415  
; CURRENT FILING DATE: 1999-06-15  
; NUMBER OF SEQ ID NOS: 13  
; SOFTWARE: PatentIn Ver. 2.0  
; SEQ ID NO 7  
; LENGTH: 39  
; TYPE: PRF  
; ORGANISM: Heloderma suspectum  
US-09-333-415-7

Query Match 95.7%; Score 200; DB 4; Length 39;  
Best Local Similarity 94.9%; Pred. No. 3..76-19;  
Matches 37; Conservative 1; Mismatches 1; Indels 0; Gaps 0;

Qy 1 HGGTFTSDLSKQMEAVRLFIEWLNGKGPSSGAPPPS 39  
Db 1 HSDGFTFTSDLSKQMEAVRLFIEWLNGKGPSSGAPPPS 39

RESULT 13  
US-09-303-016-7  
; Sequence 7, Application US/09303016  
; PATENT NO. 6429397  
; GENERAL INFORMATION:  
; APPLICANT: Coolidge, Thomas R.  
; APPLICANT: Ehlers, Mario R.W.  
; TITLE OF INVENTION: Metabolic Intervention with GLP-1 or its Biologically  
; Active Analogues to Improve the Function of the  
; Title: Ischemic and Reperfused Brain  
; FILE REFERENCE: P03660US2  
; CURRENT APPLICATION NUMBER: US/09/303,016  
; CURRENT FILING DATE: 1999-04-30  
; NUMBER OF SEQ ID NOS: 13  
; SOFTWARE: PatentIn Ver. 2.0  
; SEQ ID NO 7  
; LENGTH: 39  
; TYPE: PRF  
; ORGANISM: Heloderma suspectum  
US-09-303-016-7

RESULT 14  
 US-09-623-618B-19  
 Query Match 95.7%; Score 200; DB 4; Length 39;  
 Best Local Similarity 94.9%; Pred. No. 3.7e-19;  
 Matches 37; Conservative 1; Mismatches 1; Indels 0; Gaps 0;  
 TYPE: PRT  
 SOFTWARE: FastSBQ for Windows Version 4.0  
 SEQ ID NO: 33  
 LENGTH: 40  
 SEQ ID NO: 33  
 LENGTH: 40  
 TYPE: PRT  
 FEATURE:  
 ORGANISM: Artificial Sequence  
 OTHER INFORMATION: Description of Artificial Sequence: Synthetic  
 NAME/KEY: MOD\_RES  
 LOCATION: 40  
 OTHER INFORMATION: Xaa represents Lys(E-MPA)-NH2-5TFA and where "E" represents B1  
 US-09-623-618B-33

Query Match 95.7%; Score 200; DB 4; Length 40;  
 Best Local Similarity 94.9%; Pred. No. 3.9e-19;  
 Matches 37; Conservative 1; Mismatches 1; Indels 0; Gaps 0;  
 QY 1 HGEGRFTSDLSKQMEBEAVRLFIEWLKNGPSSGAPPS 39  
 DB 1 HSDGTRFTSDLSKQMEBEAVRLFIEWLKNGPSSGAPPS 39

RESULT 15  
 US-09-623-618B-19  
 Sequence 19, Application US/09623618B  
 Patent No. 6329336  
 GENERAL INFORMATION:  
 APPLICANT: Bridor, Dominique P.  
 APPLICANT: L'Archeveque, Benoit  
 APPLICANT: Ezrin, Alan M.  
 APPLICANT: Holmes, Darren L.  
 APPLICANT: Leblanc, Anouk  
 APPLICANT: St. Pierre, Serge  
 TITLE OF INVENTION: LONG LASTING INSULINOTROPIC PEPTIDES  
 FILE REFERENCE: 500862001620  
 CURRENT FILING DATE: 2000-09-05  
 PRIOR APPLICATION NUMBER: PCT/US00/13563  
 PRIOR FILING DATE: 2000-05-17  
 PRIOR APPLICATION NUMBER: 60/159,783  
 PRIOR FILING DATE: 1999-10-15  
 PRIOR APPLICATION NUMBER: 60/134,406  
 PRIOR FILING DATE: 1999-05-17  
 NUMBER OF SEQ ID NOS: 35  
 SEQ ID NO: 35  
 SOFTWARE: FastSBQ for Windows Version 4.0  
 LENGTH: 40  
 TYPE: PRT  
 ORGANISM: Artificial Sequence  
 FEATURE:  
 OTHER INFORMATION: Description of Artificial Sequence: Synthetic  
 SEQ ID NO: 33  
 LENGTH: 40  
 SEQ ID NO: 33  
 LENGTH: 40  
 TYPE: PRT  
 FEATURE:  
 OTHER INFORMATION: Description of Artificial Sequence: Synthetic  
 SEQ ID NO: 33  
 LENGTH: 40  
 SEQ ID NO: 33  
 LENGTH: 40  
 TYPE: PRT  
 FEATURE:  
 OTHER INFORMATION: Description of Artificial Sequence: Synthetic  
 NAME/KEY: MOD\_RES  
 LOCATION: 40  
 OTHER INFORMATION: Xaa represents Lys(E-MPA)-NH2-5TFA and where "E" represents B1  
 US-09-623-618B-33

Query Match 95.7%; Score 200; DB 4; Length 40;  
 Best Local Similarity 94.9%; Pred. No. 3.9e-19;  
 Matches 37; Conservative 1; Mismatches 1; Indels 0; Gaps 0;  
 QY 1 HGEGRFTSDLSKQMEBEAVRLFIEWLKNGPSSGAPPS 39  
 DB 1 HSDGTRFTSDLSKQMEBEAVRLFIEWLKNGPSSGAPPS 39

GenCore version 5.1.3  
Copyright (c) 1993 - 2003 Compugen Ltd.

OM protein - protein search, using sw model

Run on: 4 February 13, 2003, 17:10:53 ; Search time 11 Seconds  
(without alignment(s), 90.582 Million cell updates/sec)

Title: US-09-756-690A-2  
Perfect score: 209  
Sequence: 1 HGEGETFTSDLKQMEEEAVRLFIWKLNGPSSGAPPS 39

Scoring table: BLOSUM62  
Gapop 10.0 , Gapext 0.5

Searched: 140259 seqs, 2548876 residues

Total number of hits satisfying chosen parameters: 140259

Minimum DB seq length: 0  
Maximum DB seq length: 0  
Post-processing: Minimum Match 0%  
Maximum Match 100%  
Listing first 45 summaries

Database : Published Applications/All,\*

1: /cgm2\_6/picodata/2/pubaa/US08 NEW PUB.PEP.\*  
2: /cgm2\_6/picodata/2/pubaa/PCT NEW PUB.PEP.\*  
3: /cgm2\_6/picodata/2/pubaa/US06 NEW PUB.PEP.\*  
4: /cgm2\_6/picodata/2/pubaa/US06 PUBCOMB.PEP.\*  
5: /cgm2\_6/picodata/2/pubaa/US07 NEW PUB.PEP.\*  
6: /cgm2\_6/picodata/2/pubaa/US07 PUBCOMB.PEP.\*  
7: /cgm2\_5/picodata/2/pubaa/PCRTUS.PUBCOMB.PEP.\*  
8: /cgm2\_6/picodata/2/pubaa/US08 PUBCOMB.PEP.\*  
9: /cgm2\_6/picodata/2/pubaa/US09 NEW PUB.PEP.\*  
10: /cgm2\_6/picodata/2/pubaa/US07 PUBCOMB.PEP.\*  
11: /cgm2\_6/picodata/2/pubaa/US10 NEW PUB.PEP.\*  
12: /cgm2\_6/picodata/2/pubaa/US10 PUBCOMB.PEP.\*  
13: /cgm2\_6/picodata/2/pubaa/US60 NEW PUB.PEP.\*  
14: /cgm2\_6/picodata/2/pubaa/US60 PUBCOMB.PEP.\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

## SUMMARIES

Result No.	Score	Query Match	Length	DB ID	Description
1	209	100.0	39	10 US-09-876-388-12	Sequence 12, App1
2	209	100.0	39	10 US-09-871-738-9	Sequence 9, App1
3	209	100.0	39	10 US-09-859-507-9	Sequence 9, App1
4	209	100.0	39	10 US-09-859-804-9	Sequence 9, App1
5	209	100.0	39	10 US-09-853-859-2	Sequence 9, App1
6	209	100.0	39	10 US-09-924-978-9	Sequence 9, App1
7	209	100.0	39	10 US-09-933-011B-9	Sequence 9, App1
8	209	100.0	40	10 US-09-876-388-18	Sequence 18, App1
9	209	100.0	40	10 US-09-876-388-31	Sequence 31, App1
10	209	100.0	40	10 US-09-876-388-32	Sequence 32, App1
11	208	99.5	39	10 US-09-003-859-25	Sequence 25, App1
12	206	98.6	39	10 US-09-003-859-10	Sequence 10, App1
13	206	98.6	39	10 US-09-003-859-14	Sequence 14, App1
14	206	98.6	39	10 US-09-003-859-14	Sequence 18, App1
15	206	98.6	39	10 US-09-003-859-29	Sequence 29, App1
16	205	98.1	38	10 US-09-003-859-62	Sequence 62, App1
17	205	98.1	39	10 US-09-003-860-13	Sequence 13, App1
18	205	98.1	39	10 US-09-003-859-16	Sequence 16, App1
19	205	98.1	39	10 US-09-003-859-19	Sequence 19, App1

## ALIGNMENTS

RESULT 1  
US-09-876-388-12  
Sequence 12, Application US/09876388  
Patient No. US0020049153A1  
GENERAL INFORMATION:  
/ APPLICANT: Bridon, Dominique F.  
/ APPLICANT: L'Archeveque, Benoit  
/ APPLICANT: Bzrin, Alan M.  
/ APPLICANT: Holmes, Darren L.  
/ APPLICANT: Leblanc, Anouk  
/ APPLICANT: St. Pierre, Serge  
TITLE OF INVENTION: LONG LASTING INSULINOTROPIC PEPTIDES  
FILE REFERENCE: 50086201610  
CURRENT FILING DATE: 2001-09-24  
PRIORITY APPLICATION NUMBER: 09/623,618  
PRIORITY FILING DATE: 2000-09-05  
PRIORITY APPLICATION NUMBER: PCT/US00/13563  
PRIORITY FILING DATE: 2000-05-17  
PRIORITY APPLICATION NUMBER: 60/159,783  
PRIORITY FILING DATE: 1999-10-15  
PRIORITY APPLICATION NUMBER: 60/134,406  
NUMBER OF SEQ ID NOS: 35  
/ SOFTWARE: Patentin Ver. 2.1  
/ SEQ ID NO: 12  
/ LENGTH: 39  
/ TYPE: PRT  
/ ORGANISM: Artificial Sequence  
/ FEATURE:  
/ OTHER INFORMATION: Description of Artificial Sequence: Synthetic  
/ OTHER INFORMATION: Peptide  
US-09-876-388-12

RESULT 2  
US-09-851-738-9  
Sequence 9, Application US/09851738  
; Patent No. US2002005460A1  
; GENERAL INFORMATION  
; APPLICANT: Coolidge, Thomas R.  
; APPLICANT: Ehlers, Mario R.W.  
; TITLE OF INVENTION: Ischemic and Reperfused Tissue  
; FILE REFERENCE: P03660US1  
; CURRENT APPLICATION NUMBER: US/09/851,738  
; PRIOR APPLICATION NUMBER: 09/302,596  
; PRIORITY FILING DATE: 2001-05-09  
; NUMBER OF SEQ ID NOS: 13  
; SOFTWARE: PatentIn Ver. 2.0  
; SEQ ID NO: 9  
; LENGTH: 39  
; TYPE: PRT  
; ORGANISM: Gila Monster venom  
US-09-851-738-9

Query Match 100.0%; Score 209; DB 10; Length 39;  
Best Local Similarity 100.0%; Pred. No. 9.5e-20;  
Matches 39; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
Qy 1 HGGGTFTSDLSKOMEEARLFLIEWLNGGPSSGAPPPS 39  
Db 1 HGGGTFTSDLSKOMEEARLFLIEWLNGGPSSGAPPPS 39

RESULT 3  
US-09-805-507-9  
; Sequence 9, Application US/09805507  
; Patent No. US2002008192A1  
; GENERAL INFORMATION  
; APPLICANT: COOLIDGE, THOMAS R.  
; APPLICANT: EHLLERS, MARIO  
; TITLE OF INVENTION: TREATMENT OF ACUTE CORONARY SYNDROME WITH GLP-1  
; FILE REFERENCE: 09197/0395  
; CURRENT APPLICATION NUMBER: US/09/805,507  
; PRIOR APPLICATION NUMBER: 09/859,804  
; PRIORITY FILING DATE: 2001-03-14  
; NUMBER OF SEQ ID NOS: 13  
; SOFTWARE: PatentIn Ver. 2.1  
; SEQ ID NO: 9  
; LENGTH: 39  
; TYPE: PRT  
; ORGANISM: Unknown Organism  
; FEATURE:  
; OTHER INFORMATION: Description of Unknown Organism: Exendrin 4  
US-09-805-507-9

Query Match 100.0%; Score 209; DB 10; Length 39;  
Best Local Similarity 100.0%; Pred. No. 9.5e-20;  
Matches 39; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
Qy 1 HGGGTFTSDLSKOMEEARLFLIEWLNGGPSSGAPPPS 39  
Db 1 HGGGTFTSDLSKOMEEARLFLIEWLNGGPSSGAPPPS 39

RESULT 4  
US-09-859-804-9  
; Sequence 9, Application US/09859804  
; Patent No. US20020107206A1  
; GENERAL INFORMATION  
; APPLICANT: COOLIDGE, THOMAS R.  
; APPLICANT: EHLLERS, MARIO  
; TITLE OF INVENTION: TREATMENT OF ACUTE CORONARY SYNDROME WITH GLP-1  
; FILE REFERENCE: 089187/0395

RESULT 5  
US-09-003-869-2  
Sequence 2, Application US/09003869A  
; Patent No. US00213766A1  
; GENERAL INFORMATION  
; APPLICANT: BREILEY, NIGEL ROBERT ARNOLD  
; APPLICANT: BRICKETT, KATHRYN S.  
; APPLICANT: BHAVSAR, SUNIL  
; TITLE OF INVENTION: USE OF EXENDINS AND AGONISTS THEREOF FOR  
; THE REDUCTION OF FOOD INTAKE  
; FILE REFERENCE: 231/91  
; CURRENT APPLICATION NUMBER: US/09/003,869A  
; CURRENT FILING DATE: 1998-01-07  
; EARLIER APPLICATION NUMBER: US 60/034,905  
; EARLIER FILING DATE: 1997-01-07  
; EARLIER APPLICATION NUMBER: US 60/055,404  
; EARLIER FILING DATE: 1997-08-08  
; EARLIER APPLICATION NUMBER: US 60/065,442  
; EARLIER FILING DATE: 1997-11-14  
; EARLIER APPLICATION NUMBER: US 60/066,029  
; EARLIER FILING DATE: 1997-11-14  
; NUMBER OF SEQ ID NOS: 188  
; SOFTWARE: FastSEQ for Windows Version 3.0  
; SEQ ID NO: 2  
; LENGTH: 39  
; TYPE: PRT  
; ORGANISM: Heloderma suspectum  
; FEATURE:  
; NAME/KEY: AMIDATION  
; LOCATION: (39)...(39)  
; OTHER INFORMATION: amidated Ser (Serinamide)  
US-09-003-859-2  
Query Match 100.0%; Score 209; DB 10; Length 39;  
Best Local Similarity 100.0%; Pred. No. 9.5e-20;  
Matches 39; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
Qy 1 HGGGTFTSDLSKOMEEARLFLIEWLNGGPSSGAPPPS 39  
Db 1 HGGGTFTSDLSKOMEEARLFLIEWLNGGPSSGAPPPS 39

RESULT 6  
US-09-982-978-9  
; Sequence 9, Application US/09982978  
; Patent No. US20020146105A1  
; GENERAL INFORMATION  
; APPLICANT: COOLIDGE, THOMAS R.  
; APPLICANT: EHLLERS, MARIO  
; TITLE OF INVENTION: TREATMENT OF ACUTE CORONARY SYNDROME WITH GLP-1

FILE REFERENCE: 089187/0395  
; CURRENT APPLICATION NUMBER: US/09/982,978  
; CURRENT FILING DATE: 2001-0-22  
; PRIORITY NUMBER: 0/859,804  
; PRIORITY FILING DATE: 2001-05-18  
; PRIORITY APPLICATION NUMBER: 6/205,239  
; PRIORITY FILING DATE: 2000-05-19  
; NUMBER OF SEQ ID NOS: 13  
; SOFTWARE: PatentIn Ver. 2.1  
; SEQ ID NO: 9  
; LENGTH: 39  
; TYPE: PRT  
; ORGANISM: Unknown Organism  
; FEATURE:  
; OTHER INFORMATION: Description of Unknown Organism: Exendrin 4  
US-09-982-978-9

Query Match 100.0%; Score 209; DB 10; Length 39;  
Best Local Similarity 100.0%; Pred. No. 9.5e-20;  
Matches 39; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

Qy 1 HGEGETFTSDLSKOMEERAVRLFIEWLKNGGSSGAPPS 39  
Db 1 HGEGETFTSDLSKOMEERAVRLFIEWLKNGGSSGAPPS 39

RESULT 7  
US-09-953-021B-9  
; Sequence 9, Application US/09953021B  
; Patent No. US2002014131A1  
; GENERAL INFORMATION:  
; APPLICANT: Coolidge, Thomas L.  
; APPLICANT: Ehlers, Mario R. W.  
; TITLE OF INVENTION: Metabolic Intervention with GLP-1 to Improve the Function of Ischemic Muscle Tissue  
; TITLE OF INVENTION: Reperfused Skeletal Muscle Tissue  
; FILE REFERENCE: 03/66056  
; CURRENT APPLICATION NUMBER: US/09/953,021B  
; CURRENT FILING DATE: 2001-09-11  
; PRIORITY NUMBER: 0/302,596  
; PRIORITY FILING DATE: 1999-04-30  
; NUMBER OF SEQ ID NOS: 13  
; SOFTWARE: PatentIn Ver. 2.0  
; SEQ ID NO: 9  
; LENGTH: 39  
; TYPE: PRT  
; ORGANISM: Heloderma suspectum

Query Match 100.0%; Score 209; DB 10; Length 39;  
Best Local Similarity 100.0%; Pred. No. 9.5e-20;  
Matches 39; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

Qy 1 HGEGETFTSDLSKOMEERAVRLFIEWLKNGGSSGAPPS 39  
Db 1 HGEGETFTSDLSKOMEERAVRLFIEWLKNGGSSGAPPS 39

RESULT 8  
US-09-876-388-18  
; Sequence 18, Application US/09876388  
; Patent No. US2002004153A1  
; GENERAL INFORMATION:  
; APPLICANT: Bridon, Dominique P.  
; APPLICANT: L'Archeveque, Benoit  
; APPLICANT: Ezrin, Alan M.  
; APPLICANT: Holmes, Darren L.  
; APPLICANT: Leblanc, Anouk  
; APPLICANT: St. Pierre, Serge  
; TITLE OF INVENTION: LONG LASTING INSULINOTROPIC PEPTIDES  
; FILE REFERENCE: 5/00862001610  
; CURRENT APPLICATION NUMBER: US/09/876,388  
; CURRENT FILING DATE: 2001-09-24  
; PRIORITY NUMBER: 09/623,618  
; PRIORITY FILING DATE: 2000-09-05  
; PRIORITY NUMBER: PCT/US00/13563  
; PRIORITY FILING DATE: 2000-05-17  
; PRIORITY APPLICATION NUMBER: 60/159,783  
; PRIORITY FILING DATE: 1999-10-15  
; PRIORITY NUMBER: 60/134,406  
; PRIORITY FILING DATE: 1999-05-17  
; NUMBER OF SEQ ID NOS: 35  
; SOFTWARE: PatentIn Ver. 2.1  
; SEQ ID NO: 31  
; LENGTH: 40  
; TYPE: PRT  
; ORGANISM: Artificial Sequence  
; FEATURE:  
; OTHER INFORMATION: Description of Artificial Sequence: Synthetic  
; NAME/KEY: MOD RES  
; LOCATION: 40  
; OTHER INFORMATION: Xaa represents Lys(E-NPA)-NH2-5TFA and where "E" represents Et

US-09-876-388-31  
; Sequence 31, Application US/09876388  
; Patent No. US2002004153A1  
; GENERAL INFORMATION:  
; APPLICANT: Bridon, Dominique P.  
; APPLICANT: L'Archeveque, Benoit  
; APPLICANT: Ezrin, Alan M.  
; APPLICANT: Holmes, Darren L.  
; APPLICANT: Leblanc, Anouk  
; APPLICANT: St. Pierre, Serge  
; TITLE OF INVENTION: LONG LASTING INSULINOTROPIC PEPTIDES  
; FILE REFERENCE: 5/00862001610  
; CURRENT APPLICATION NUMBER: US/09/876,388  
; CURRENT FILING DATE: 2001-09-24  
; PRIORITY NUMBER: 09/623,618  
; PRIORITY FILING DATE: 2000-09-05  
; PRIORITY NUMBER: PCT/US00/13563  
; PRIORITY FILING DATE: 2000-05-17  
; PRIORITY APPLICATION NUMBER: 60/159,783  
; PRIORITY FILING DATE: 1999-10-15  
; PRIORITY NUMBER: 60/134,406  
; PRIORITY FILING DATE: 1999-05-17  
; NUMBER OF SEQ ID NOS: 35  
; SOFTWARE: PatentIn Ver. 2.1  
; SEQ ID NO: 18  
; LENGTH: 40  
; TYPE: PRT  
; ORGANISM: Artificial Sequence  
; FEATURE:  
; OTHER INFORMATION: Description of Artificial Sequence: Synthetic  
; NAME/KEY: MOD RES  
; LOCATION: 40  
; OTHER INFORMATION: Xaa represents Lys(E-NPA)-NH2-5TFA and where "E" represents Et

US-09-876-388-31  
; Sequence 31, Application US/09876388  
; Patent No. US2002004153A1  
; GENERAL INFORMATION:  
; APPLICANT: Bridon, Dominique P.  
; APPLICANT: L'Archeveque, Benoit  
; APPLICANT: Ezrin, Alan M.  
; APPLICANT: Holmes, Darren L.  
; APPLICANT: Leblanc, Anouk  
; APPLICANT: St. Pierre, Serge  
; TITLE OF INVENTION: LONG LASTING INSULINOTROPIC PEPTIDES  
; FILE REFERENCE: 5/00862001610  
; CURRENT APPLICATION NUMBER: US/09/876,388  
; CURRENT FILING DATE: 2001-09-24  
; PRIORITY NUMBER: 09/623,618  
; PRIORITY FILING DATE: 2000-09-05  
; PRIORITY NUMBER: PCT/US00/13563  
; PRIORITY FILING DATE: 2000-05-17  
; PRIORITY APPLICATION NUMBER: 60/159,783  
; PRIORITY FILING DATE: 1999-10-15  
; PRIORITY NUMBER: 60/134,406  
; PRIORITY FILING DATE: 1999-05-17  
; NUMBER OF SEQ ID NOS: 35  
; SOFTWARE: PatentIn Ver. 2.1  
; SEQ ID NO: 31  
; LENGTH: 40  
; TYPE: PRT  
; ORGANISM: Artificial Sequence  
; FEATURE:  
; OTHER INFORMATION: Description of Artificial Sequence: Synthetic  
; NAME/KEY: MOD RES  
; LOCATION: 40  
; OTHER INFORMATION: Xaa represents Lys(E-NPA)-NH2-5TFA and where "E" represents Et

RESULT 10  
 US-09-876-3-88-32  
 Sequence 32, Application US/09876388  
 Patent No. US2002009153A1  
 GENERAL INFORMATION:  
 APPLICANT: Bridon, Dominique P.  
 APPLICANT: L'Archeveque, Benoit  
 APPLICANT: Ezrin, Alan M.  
 APPLICANT: Holmes, Darren L.  
 APPLICANT: Leblanc, Anouk  
 TITLE OF INVENTION: LONG LASTING INSULINOTROPIC PEPTIDES  
 FILE REFERENCE: 500862001610  
 CURRENT APPLICATION NUMBER: US/09/876, 388  
 CURRENT FILING DATE: 2001-09-24  
 PRIOR FILING DATE: 2000-09-05  
 PRIOR APPLICATION NUMBER: PCT/US00/13565  
 PRIOR FILING DATE: 2000-05-17  
 PRIOR APPLICATION NUMBER: PCT/US00/134, 406  
 PRIOR FILING DATE: 1999-10-15  
 PRIOR APPLICATION NUMBER: 60/159, 783  
 SEQ ID NO: 32  
 LENGTH: 40  
 TYPE: PRT  
 FEATURE:  
 OTHER INFORMATION: Description of Artificial Sequence: Synthetic  
 OTHER INFORMATION: Peptide  
 NAME/KEY: MOD\_RES  
 LOCATION: 40  
 OTHER INFORMATION: Xaa represents Lys (B-ABEA-ABEA-MPA) -NH2-5TFA and where "E" represents  
 US-09-876-3-88-32  
 - Query Match 100.0%; Score 209; DB 10; Length 40;  
 - Best Local Similarity 100.0%; Pred. No. 9. Be-20;  
 - Matches 39; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
 Qy 1 HGEGETPDSLQKMEEARLFIENLKGCGPSSGAPPPS 39  
 Db 1 HGEGETPDSLQKMEEARLFIENLKGCGPSSGAPPPS 39

RESULT 11  
 US-09-003-869-25  
 Sequence 25, Application US/09003869A  
 Patent No. US2002013766A1  
 GENERAL INFORMATION:  
 APPLICANT: BEELEY, NIGEL ROBERT ARNOLD  
 APPLICANT: PRICKETT, KATHRYN S.  
 APPLICANT: BEAVSAR, SUNIL  
 APPLICANT: BRASSAR, SUNIL  
 TITLE OF INVENTION: USE OF EXENDINS AND AGONISTS THEREOF FOR  
 THE REDUCTION OF FOOD INTAKE  
 FILE REFERENCE: 231/181  
 CURRENT APPLICATION NUMBER: US/09/003, 869A  
 CURRENT FILING DATE: 1998-01-07  
 EARLIER APPLICATION NUMBER: US 60/034, 905  
 EARLIER FILING DATE: 1997-01-07  
 EARLIER APPLICATION NUMBER: US 60/055, 404  
 EARLIER FILING DATE: 1997-08-08  
 EARLIER APPLICATION NUMBER: US 60/065, 442  
 EARLIER FILING DATE: 1997-11-14  
 EARLIER APPLICATION NUMBER: US 60/066, 029  
 EARLIER FILING DATE: 1997-11-14  
 NUMBER OF SEQ ID NOS: 188  
 SEQ ID NO 10  
 LENGTH: 39  
 TYPE: PRT  
 ORGANISM: Artificial Sequence  
 FEATURE:  
 OTHER INFORMATION: Artificial Sequence  
 OTHER INFORMATION: Compound  
 FEATURE:  
 NAME/KEY: AMIDATION  
 LOCATION: (39)  
 OTHER INFORMATION: amidated Ser (Serinamide)  
 US-09-003-869-10  
 - Query Match 99.5%; Score 208; DB 10; Length 39;  
 - Best Local Similarity 97.4%; Pred. No. 1.3e-19;  
 - Matches 38; Conservative 1; Mismatches 0; Indels 0; Gaps 0;  
 Qy 1 HGBGFTPSDSLQKMEEARLFIENLKGCGPSSGAPPPS 39  
 Db 1 HGGGFTPSDSLQKMEEARLFIENLKGCGPSSGAPPPS 39

RESULT 12  
 US-09-003-869-10  
 Sequence 10, Application US/09003869A  
 Patent No. US2002013766A1  
 GENERAL INFORMATION:  
 APPLICANT: BEELEY, NIGEL ROBERT ARNOLD  
 APPLICANT: PRICKETT, KATHRYN S.  
 APPLICANT: BEAVSAR, SUNIL  
 APPLICANT: BRASSAR, SUNIL  
 TITLE OF INVENTION: USE OF EXENDINS AND AGONISTS THEREOF FOR  
 THE REDUCTION OF FOOD INTAKE  
 FILE REFERENCE: 231/181  
 CURRENT APPLICATION NUMBER: US/09/003, 869A  
 CURRENT FILING DATE: 1998-01-07  
 EARLIER APPLICATION NUMBER: US 60/034, 905  
 EARLIER FILING DATE: 1997-01-07  
 EARLIER APPLICATION NUMBER: US 60/055, 404  
 EARLIER FILING DATE: 1997-08-08  
 EARLIER APPLICATION NUMBER: US 60/065, 442  
 EARLIER FILING DATE: 1997-11-14  
 EARLIER APPLICATION NUMBER: US 60/066, 029  
 EARLIER FILING DATE: 1997-11-14  
 NUMBER OF SEQ ID NOS: 188  
 SEQ ID NO 10  
 LENGTH: 39  
 TYPE: PRT  
 ORGANISM: Artificial Sequence  
 FEATURE:  
 OTHER INFORMATION: Artificial Sequence  
 OTHER INFORMATION: Compound  
 FEATURE:  
 NAME/KEY: AMIDATION  
 LOCATION: (39)  
 OTHER INFORMATION: amidated Ser (Serinamide)  
 US-09-003-869-10  
 - Query Match 98.6%; Score 206; DB 10; Length 39;  
 - Best Local Similarity 97.4%; Pred. No. 2.3e-19;  
 - Matches 38; Conservative 1; Mismatches 0; Indels 0; Gaps 0;  
 Qy 1 HGBGFTPSDSLQKMEEARLFIENLKGCGPSSGAPPPS 39  
 Db 1 HGGGFTPSDSLQKMEEARLFIENLKGCGPSSGAPPPS 39

RESULT 13  
 US-09-003-869-14  
 Sequence 14, Application US/09003869A  
 Patent No. US2002013766A1  
 GENERAL INFORMATION:  
 APPLICANT: BEELEY, NIGEL ROBERT ARNOLD  
 APPLICANT: PRICKETT, KATHRYN S.  
 APPLICANT: BEAVSAR, SUNIL  
 APPLICANT: BRASSAR, SUNIL  
 TITLE OF INVENTION: USE OF EXENDINS AND AGONISTS THEREOF FOR  
 THE REDUCTION OF FOOD INTAKE  
 FILE REFERENCE: 231/181  
 CURRENT APPLICATION NUMBER: US/09/003, 869A  
 CURRENT FILING DATE: 1998-01-07  
 EARLIER APPLICATION NUMBER: US 60/065, 442  
 EARLIER FILING DATE: 1997-11-14  
 EARLIER APPLICATION NUMBER: US 60/055, 404  
 EARLIER FILING DATE: 1997-08-08  
 NUMBER OF SEQ ID NOS: 188  
 SOFTWARE: PastSeq For Windows Version 3.0  
 SEQ ID NO: 25  
 LENGTH: 39  
 TYPE: PRT  
 ORGANISM: Artificial Sequence

FILE REFERENCE: 231/181  
 CURRENT APPLICATION NUMBER: US/09/003, 869A  
 CURRENT FILING DATE: 1998-01-07  
 EARLIER APPLICATION NUMBER: US 60/034, 905  
 EARLIER FILING DATE: 1997-01-07  
 EARLIER APPLICATION NUMBER: US 60/055, 104  
 EARLIER FILING DATE: 1997-08-08  
 EARLIER APPLICATION NUMBER: US 60/065, 442  
 EARLIER FILING DATE: 1997-11-14  
 EARLIER APPLICATION NUMBER: US 60/066, 029  
 EARLIER FILING DATE: 1997-11-14  
 NUMBER OF SEQ ID NOS: 188  
 SOFTWARE: FastSEQ For Windows Version 3.0  
 SEQ ID NO: 14  
 TYPE: PRT  
 ORGANISM: Artificial Sequence  
 FEATURE:  
 OTHER INFORMATION: artificially synthesized sequence of novel exendin agonist  
 OTHER INFORMATION: compound  
 FEATURE:  
 NAME/KEY: AMIDATION  
 LOCATION: (39) .. (39)  
 OTHER INFORMATION: amidated Ser (Serinamide)  
 US-09-003-869-14

Query Match 98.6%; Score 206; DB 10; Length 39;  
 Best Local Similarity 97.4%; Pred. No. 2.3e-19;  
 Matches 38; Conservative 1; Mismatches 0;  
 Indels 0; Gaps 0;

Qy 1 HGGCTTSDLSKQMEEARLFWLNGGPGSGAPPS 39  
 Db 1 HGGCTTSDLSKQMEEARLFWLNGGPGSGAPPS 39

RESULT 14  
 US-09-003-869-18  
 Sequence 18. Application US/09/003, 869A  
 Patent No. US20033766A1  
 GENERAL INFORMATION:  
 APPLICANT: BEELEY, NIGEL ROBERT ARNOLD  
 APPLICANT: PRICKETT, KATHRYN S.  
 APPLICANT: BHAVSAR, SUNIL  
 TITLE OF INVENTION: USE OF EXENDINS AND AGONISTS THEREOF FOR  
 TITLE OF INVENTION: THE REDUCTION OF FOOD INTAKE  
 FILE REFERENCE: 231/181  
 CURRENT APPLICATION NUMBER: US/09/003, 869A  
 CURRENT FILING DATE: 1998-01-07  
 EARLIER APPLICATION NUMBER: US 60/034, 905  
 EARLIER FILING DATE: 1997-01-07  
 EARLIER APPLICATION NUMBER: US 60/065, 442  
 EARLIER FILING DATE: 1997-08-08  
 EARLIER APPLICATION NUMBER: US 60/066, 029  
 EARLIER FILING DATE: 1997-11-14  
 NUMBER OF SEQ ID NOS: 188  
 SOFTWARE: FastSEQ for Windows Version 3.0  
 SEQ ID NO: 29  
 LENGTH: 39  
 TYPE: PRT  
 ORGANISM: Artificial Sequence  
 FEATURE:  
 OTHER INFORMATION: artificially synthesized sequence of novel exendin agonist  
 OTHER INFORMATION: Compound  
 FEATURE:  
 NAME/KEY: AMIDATION  
 LOCATION: (39) .. (39)  
 OTHER INFORMATION: amidated Ser (Serinamide)  
 US-09-003-869-29

Query Match 98.6%; Score 206; DB 10; Length 39;  
 Best Local Similarity 97.4%; Pred. No. 2.3e-19;  
 Matches 38; Conservative 1; Mismatches 0;  
 Indels 0; Gaps 0;

Qy 1 HGGCTTSDLSKQMEEARLFWLNGGPGSGAPPS 39  
 Db 1 HGGCTTSDLSKQMEEARLFWLNGGPGSGAPPS 39

RESULT 15  
 US-09-003-869-29  
 Sequence 29. Application US/09/003, 869A  
 Patent No. US20033766A1  
 GENERAL INFORMATION:  
 APPLICANT: BEELEY, NIGEL ROBERT ARNOLD  
 APPLICANT: PRICKETT, KATHRYN S.  
 APPLICANT: BHAVSAR, SUNIL  
 TITLE OF INVENTION: USE OF EXENDINS AND AGONISTS THEREOF FOR  
 TITLE OF INVENTION: THE REDUCTION OF FOOD INTAKE  
 FILE REFERENCE: 231/181  
 CURRENT APPLICATION NUMBER: US/09/003, 869A  
 CURRENT FILING DATE: 1998-01-07  
 EARLIER APPLICATION NUMBER: US 60/034, 905  
 EARLIER FILING DATE: 1997-01-07  
 EARLIER APPLICATION NUMBER: US 60/065, 442  
 EARLIER FILING DATE: 1997-08-08  
 EARLIER APPLICATION NUMBER: US 60/066, 029  
 EARLIER FILING DATE: 1997-11-14  
 NUMBER OF SEQ ID NOS: 188  
 SOFTWARE: FastSEQ for Windows Version 3.0  
 SEQ ID NO: 29  
 LENGTH: 39  
 TYPE: PRT  
 ORGANISM: Artificial Sequence  
 FEATURE:  
 OTHER INFORMATION: artificially synthesized sequence of novel exendin agonist  
 OTHER INFORMATION: compound  
 FEATURE:  
 NAME/KEY: AMIDATION  
 LOCATION: (39) .. (39)  
 OTHER INFORMATION: amidated Ser (Serinamide)  
 US-09-003-869-29

Query Match 98.6%; Score 206; DB 10; Length 39;  
 Best Local Similarity 97.4%; Pred. No. 2.3e-19;  
 Matches 38; Conservative 1; Mismatches 0;  
 Indels 0; Gaps 0;

Qy 1 HGGCTTSDLSKQMEEARLFWLNGGPGSGAPPS 39  
 Db 1 HGGCTTSDLSKQMEEARLFWLNGGPGSGAPPS 39

Search completed: February 13, 2003, 17:13:45  
 Job time : 22 secs

Query Match 98.6%; Score 206; DB 10; Length 39;  
 Best Local Similarity 97.4%; Pred. No. 2.3e-19;  
 Matches 38; Conservative 1; Mismatches 0;  
 Indels 0; Gaps 0;

Qy 1 HGGCTTSDLSKQMEEARLFWLNGGPGSGAPPS 39  
 Db 1 HGGCTTSDLSKQMEEARLFWLNGGPGSGAPPS 39

Query Match 98.6%; Score 206; DB 10; Length 39;  
 Best Local Similarity 97.4%; Pred. No. 2.3e-19;

GenCore version 5.1.3  
Copyright (c) 1993 - 2003 Compugen Ltd.

OM protein - protein search, using sw model

Run on: 4 February 13, 2003, 17:08:43 ; Search time 47 Seconds  
(without alignments)

79.771 Million cell updates/sec

Title: US-09-756-690a-2  
Perfect score: 209  
Sequence: 1 HGEGETFTSDLSKQMEBEAVRLFIKNGPSSGAPPS 39

Scoring table: BLOSUM62  
Gapop 10.0 , Gapext 0.5

Searched: 283224 seqs, 9613422 residues

Total number of hits satisfying chosen parameters: 283224

Minimum DB seq length: 0  
Maximum DB seq length: 2000000000Post-processing: Minimum Match 0%  
Maximum Match 100%  
Listing first 45 summaries

Database : PIR:73:\*

1: pir1:\*

2: pir2:\*

3: pir3:\*

4: pir4:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

## SUMMARIES

Result No.	Score	Query	Match	Length	DB ID	Description
1	209	100.0	1	HWGH4G		exendin-4 - Gila m
2	200	95.7	39	1	HWGH3Z	exendin-3 - Mexica
3	99	47.4	31	2	S44472	Glucagon G2 - Nort
4	97	46.4	31	2	S44471	Glucagon G1 Nort
5	97	46.4	101	1	GCFGB	Glucagon Precursor
6	94	45.0	63	1	GCIDC	Glucagon Precursor
7	92	44.0	30	2	B61125	Glucagon-like pept
8	92	44.0	30	2	C61125	Glucagon-like pept
9	91	43.5	30	2	S44473	Glucagon-like pept
10	90.5	43.3	178	2	S44478	Glucagon I precurso
11	89	42.6	72	1	GCGXA	Glucagon Precursor
12	88	42.1	66	1	151093	Glucagon - chinook
13	88	42.1	151	1	GCCH	Glucagon Precursor
14	88	42.1	178	2	151057	Glucagon II precurso
15	88	42.1	206	2	151301	Glucagon - chick
16	87	41.6	29	1	GCDF	Glucagon - smaller
17	87	41.6	158	1	GCPG	Glucagon precursor
18	87	41.6	180	1	GCNU	Glucagon precursor
19	87	41.6	180	1	GCGP	Glucagon precursor
20	87	41.6	180	1	GRTRDU	Glucagon precursor
21	87	41.6	180	1	GCRT	Glucagon precursor
22	87	41.6	180	1	GCYI	Glucagon precursor
23	87	41.6	180	1	GCBO	Glucagon precursor
24	87	41.6	180	2	A57294	Glucagon precursor
25	86	41.1	122	1	GCAP2	Glucagon 2 precurso
26	84	40.2	29	1	GCPL	Glucagon - Europea
27	84	40.2	29	2	S07211	Glucagon - marbled
28	84	40.2	29	2	A61135	Glucagon - bigeye
29	84	40.2	67	1	GCFS	Glucagon precursor

## ALIGNMENTS

RESULT 1  
HWGH4G  
exendin-4 - Gila monster  
C;Species: Heloderma suspectum (Gila monster)  
C;Date: 31-Mar-1993 #sequence\_revision 31-Mar-1993 #text\_change 21-Nov-1997  
C;Accession: A42486  
R;Eng, J.; Kleiman, W.A.; Singh, L.; Singh, G.; Raufman, J.P.  
J. Biol. Chem. 267, 7402-7405, 1992  
A;Title: Isolation and characterization of exendin-4, an exendin-3 analogue, from Heloderma suspectum  
A;Reference number: A42486; PMID:1313797  
A;Accession: A42486  
A;Molecule type: protein  
A;Residues: 1-39 <ENG>  
C;Superfamily: Glucagon  
C;Keywords: amidated carboxyl end; duplication; venom  
E;39/Modified site: amidated carboxyl end (Ser) #status experimental

Query Match 100.0% ; Score 209 ; DB 1; Length 39;  
Best Local Similarity 100.0% ; Pred. No. 2.6e-19;  
Matches 39 ; Conservative 0 ; Mismatches 0 ; Indels 0 ; Gaps 0 ;

Query Match 100.0% ; Score 209 ; DB 1; Length 39;  
Best Local Similarity 100.0% ; Pred. No. 2.6e-19;  
Matches 39 ; Conservative 0 ; Mismatches 0 ; Indels 0 ; Gaps 0 ;

RESULT 2  
HWGH4Z  
exendin-3 - Mexican beaded lizard  
C;Species: Heloderma horridum (Mexican beaded lizard)  
C;Date: 31-Mar-1993 #sequence\_revision 31-Mar-1993 #text\_change 21-Nov-1997  
C;Accession: A23674  
R;Eng, J.; Andrews, P.C.; Kleinman, W.A.; Singh, L.; Raufman, J.P.  
J. Biol. Chem. 265, 20259-20262, 1990  
A;Title: Purification and structure of exendin-3, a new pancreatic secretagogue isolat  
A;Reference number: A23674; PMID:1700785  
A;Accession: A23674  
A;Molecule type: protein  
A;Residues: 1-39 <ENG>  
C;Comment: Exendins are venom components that are thought to bind to receptors for v  
g in secretion of amylase.  
C;Superfamily: Glucagon  
C;Keywords: amidated carboxyl end; duplication; secretagogue; venom  
E;39/Modified site: amidated carboxyl end (Ser) #status experimental

Query Match 95.7% ; Score 200 ; DB 1; Length 39;  
Best Local Similarity 94.9% ; Pred. No. 3.4e-18;  
Matches 37 ; Conservative 1 ; Mismatches 1 ; Indels 0 ; Gaps 0 ;

Query Match 1 HEGETFTSDLSKQMEBEAVRLFIKNGPSSGAPPS 39

Db 1 HSDGFTSDLSKQMEEAIRLIEWLNGPSSGAPPSS 39

RESULT 3

Db 1 HSDGFTSDLSKQMEEAIRLIEWLNGPSSGAPPSS 39

Glucagon G2 - North American paddlefish (Polyodon spathula)

C:Species: Polyodon spathula

C:Accession: S44472

C:Superfamily: 19-Mar-1997 #sequence\_revision 12-Dec-1997 #text\_change 07-May-1999

C:Accession: S44472

C:Title: Characterization of insulin and proglucagon-derived peptides from a phylogenetic perspective

A:Reference number: S44467; MUID:94271144; PMID:8002937

A:Accession: S44472

A:Molecule type: protein

A:Residues: 1-31 <NGD>

A:Note: the sequence from Fig. 3 is inconsistent with that from Fig. 5 in having 29-Glu

C:Keywords: Glucagon

C:Superfamily: carbohydrate metabolism; duplication; hormone; pancreas

F:1-31/Product: Glucagon G2 #status predicted <GCN>

Query Match 47.4% Score 99; DB 2; Length 31;

Best Local Similarity 55.2%; Pred. No. 9.4e-06; Indels 0; Gaps 0;

Matches 16; Conservative 7; Mismatches 6; Indels 0; Gaps 0;

Db 1 HSQGMFTNDYSKYLDEKAEKFWLNGK 29

RESULT 4

Db 1 HSGCFTSDLSKQMEEAIRLIEWLNGK 29

Glucagon G1 - North American paddlefish (Polyodon spathula)

C:Species: Polyodon spathula

C:Accession: S44471

C:Superfamily: 18-Sep-1997 #sequence\_revision 18-Sep-1997 #text\_change 07-May-1999

C:Accession: S44471

C:Title: Characterization of insulin and proglucagon-derived peptides from a phylogenetic perspective

A:Reference number: S44467; MUID:94271144; PMID:8002937

A:Accession: S44471

A:Molecule type: protein

A:Residues: 1-31 <NGD>

A:Experimental source: pancreas

C:Superfamily: Glucagon

C:Keywords: carbohydrate metabolism; duplication; hormone; pancreas

F:1-31/Product: Glucagon G1 #status predicted <HAT>

Query Match 46.4% Score 97; DB 2; Length 31;

Best Local Similarity 55.2%; Pred. No. 1.7e-05; Indels 0; Gaps 0;

Matches 16; Conservative 6; Mismatches 7; Indels 0; Gaps 0;

RESULT 5

Db 1 HSGCFTSDLSKQMEEAIRLIEWLNGK 29

Glucagon precursor - bullfrog (fragments)

N:Contains: Glucagon; Glucagon-16 (oxynormodulin)

C:Species: Rana catesbeiana (bullfrog)

C:Accession: B28091

C:Date: 31-Mar-1993 #sequence\_revision 31-Mar-1993 #text\_change 20-Mar-1998

C:Accession: B28091

C:Title: Isolation of peptide hormones from the pancreas of the bullfrog (Rana catesbeiana)

A:Reference number: A92730; MUID:3260236

A:Molecule type: protein

A:Residues: 1-36 <PO2>

A:Accession: C28091

Query Match 44.0% Score 92; DB 2; Length 30;

Best Local Similarity 48.3%; Pred. No. 6.7e-05; Indels 0; Gaps 0;

Matches 14; Conservative 8; Mismatches 7; Indels 0; Gaps 0;

Qy 1 HGEGTFTSDLSKQMBEEAVRLFIEWLKNG 29  
 Db 1 HAEGTFTSDVSYLQDQAKEYFSQLKNG 29

RESULT 8

Glucagon-like peptide - European eel  
 C;Species: Anguilla anguilla (European eel)  
 C;Accession: C61125  
 C;Date: 10-Mar-1994 #sequence\_revision 10-Mar-1994 #text\_change 21-Nov-1997  
 R;Conlon, J.M.; Andrews, P.C.; Thim, L.; Moon, T.W.  
 Gen. Comp. Endocrinol. 82, 23-32, 1991  
 A;Title: The primary structure of glucagon-like peptide but not insulin has been conserv  
 A;Reference number: A61125; MUID:91340688; PMID:1874385  
 A;Accession: C61125  
 A;Molecule type: Protein  
 A;Residues: 1-30 <CON>  
 C;Superfamily: Glucagon  
 C;Keywords: amidated carboxyl end; duplication F;1-30/Product: Glucagon-like peptide #status experimental <GLP>  
 F;30/Modified site: amidated carboxyl end (Arg) #status experimental

Query Match 44.0%; Score 92; DB 2; Length 30;  
 Best Local Similarity 48.3%; Pred. No. 6.7e-05;  
 Matches 14; Conservative 8; Mismatches 7; Indels 0; Gaps 0;

Qy 1 HGEGTFTSDLSKQMBEEAVRLFIEWLKNG 29  
 Db 1 HAEGTFTSDVSYLQDQAKEYFSQLKNG 29

RESULT 9

Glucagon-like peptide - North American paddlefish (Polyodon spathula)  
 C;Species: Polyodon spathula  
 C;Accession: S44473  
 C;Date: 18-Sep-1997 #sequence\_revision 18-Sep-1997 #text\_change 07-May-1999  
 R;Nguyen, T.M.; Mommisen, T.P.; Mims, S.M.; Conlon, J.M.  
 Biochem. J. 300, 339-345, 1994  
 A;Title: Characterization of insulin and proglucagon-derived peptides from a phylogenetic  
 A;Reference number: S44467; MUID:94271144; PMID:8002937  
 A;Accession: S44473  
 A;Molecule type: protein  
 A;Residues: 1-30 <NG>  
 C;Superfamily: Glucagon  
 C;Keywords: duplication; hormone; pancreas  
 F;1-30/Product: glucagon-like peptide #status predicted <MAT>

Query Match 43.5%; Score 91; DB 2; Length 30;  
 Best Local Similarity 55.2%; Pred. No. 8.9e-05;  
 Matches 16; Conservative 5; Mismatches 8; Indels 0; Gaps 0;

Qy 1 HGEGTFTSDLSKQMBEEAVRLFIEWLKNG 29  
 Db 1 HAEGTFTSDVSYLQDQAKEYFSQLKNG 29

RESULT 10

Glucagon I precursor - rainbow trout  
 C;Species: Oncorhynchus mykiss  
 C;Date: 13-Sep-1996 #sequence\_revision 13-Sep-1996 #text\_change 16-Jul-1999  
 C;Accession: I51058; I51299; I51056; 151037; 151300  
 R;Irwin, D.M.; Wong, J.  
 Mol. Endocrinol. 9, 267-277, 1995  
 A;Title: Trout and chicken proglucagon: alternative splicing generates mRNA transcripts  
 A;Reference number: A55895; MUID:776767  
 A;Accession: I51058  
 A;Status: preliminary; translated from GB/EMBL/DBJ  
 A;Molecule type: mRNA  
 A;Residues: 1-178 <IRW>  
 A;Cross-references: EMBL:U19917; NID:9736364; PMID:9736369.1; PIDN: AAC59669.1; GB:S78475; N  
 C;Date: 13-Sep-1996 #sequence\_revision 13-Sep-1996 #text\_change 16-Jul-1999

RESULT 12

Glucagon - Chinook salmon (fragment)  
 C;Species: Oncorhynchus tshawytscha (chinook salmon)  
 C;Date: 13-Sep-1996 #sequence\_revision 13-Sep-1996 #text\_change 16-Jul-1999

Qy 1 HGEGTFTSDLSKQMBEEAVRLFIEWLKNG 29  
 Db 39 HADGTYTSDVSYLQDQAKEYFSQLKNG 67

RESULT 151058

Glucagon I precursor - rainbow trout  
 C;Species: Oncorhynchus mykiss  
 C;Date: 13-Sep-1996 #sequence\_revision 13-Sep-1996 #text\_change 16-Jul-1999  
 C;Accession: I51058; I51299; I51056; 151037; 151300  
 R;Irwin, D.M.; Wong, J.  
 Mol. Endocrinol. 9, 267-277, 1995  
 A;Title: Trout and chicken proglucagon: alternative splicing generates mRNA transcripts  
 A;Reference number: A55895; MUID:776767  
 A;Accession: I51058  
 A;Status: preliminary; translated from GB/EMBL/DBJ  
 A;Molecule type: mRNA  
 A;Residues: 1-178 <IRW>  
 A;Cross-references: EMBL:U19917; NID:9736364; PMID:9736369.1; PIDN: AAC59669.1; GB:S78475; N  
 C;Date: 13-Sep-1996 #sequence\_revision 13-Sep-1996 #text\_change 16-Jul-1999

C;Accession: 151093  
 R;Irwin, D.M.; Wong, J.  
 Mol. Endocrinol. 9, 267-277, 1995  
 A;Title: Trout and chicken proglucagon: alternative splicing generates mRNA transcripts  
 A;Reference number: A55895; MUID:95295739; PMID:776976  
 A;Accession: 151093  
 A;Status: preliminary; translated from GB/EMBL/DDBJ  
 A;Molecule type: mRNA  
 A;Residues: 1-66 <IRW>  
 C;Cross-references: EMBL:U19920; NID:9736366; PID:973670.1; PID:97363367  
 C;Superfamily: glucagon  
 C;Keywords: duplication  
 C;Keywords: duplication  
 Query Match 42.1%; Score 88; DB 2; Length 66;  
 Best Local Similarity 44.8%; Pred. No. 0.00049;  
 Matches 13; Conservative 10; Missmatches 6; Indels 0; Gaps 0;  
 Qy 1 HGGTFTSDLSKQMEEEAVRLFIEWLKG 29  
 Db 33 HADGTTSDVSTYLDQQAQDFVSMUKSG 61

RESULT 13

GCCH  
 Glucagon precursor - chicken  
 N;Contains: Glucagon; glucagon-like peptide 1  
 C;Species: Gallus gallus (chicken)  
 C;Accession: S09992; A92189; A92189; A05836; A01542  
 R;Hasegawa, S.; Terazono, K.; Nata, K.; Takada, T.; Yamamoto, H.; Okamoto, H.  
 PESS Lett., 264, 117-120, 1990  
 A;Title: Nucleotide sequence determination of chicken glucagon precursor cDNA. Chicken  
 A;Reference number: S09992; MUID:90249492; PMID:2338135

A;Accession: S09992  
 A;Molecule type: mRNA  
 A;Residues: 1-151 <IRW>  
 A;Cross-references: EMBL:U07539; NID:963749; PID:9468827.1; PID:963750  
 R;Pollock, H.G.; Kimmel, J.R.  
 J. Biol. Chem. 250, 9371-9380, 1975  
 A;Title: Chicken glucagon. Isolation and amino acid sequence studies.  
 A;Reference number: A92189; MUID:70069271; PMID:1124290  
 A;Accession: A92189  
 A;Molecule type: protein  
 A;Residues: 55-83 <POL>  
 C;Superfamily: glucagon  
 C;Keywords: amidated carboxyl end; carbohydrate metabolism; duplication; hormone; pancreas  
 F;1-22/Domain: signal sequence  
 F;23-151/Product: proglucagon #status predicted <PGC>  
 F;55-83/Product: glucagon #status experimental <GCN>  
 F;118-147/Product: glucagon-like peptide 1 #status predicted <GL1>  
 F;147/Modifid site: amidated carboxyl end (Arg) tandem in mature form from following 91

Query Match 42.1%; Score 88; DB 1; Length 151;  
 Best Local Similarity 51.7%; Pred. No. 0.0012; 6; Indels 0; Gaps 0;  
 Matches 15; Conservative 6; Missmatches 6; Indels 0; Gaps 0;

Qy 1 HGGTFTSDLSKQMEEEAVRLFIEWLKG 29  
 Db 118 HADGTTSDVSTYLDQQAQDFVSMUKSG 146

RESULT 14

151057  
 Glucagon II precursor - rainbow trout  
 C;Species: Oncorhynchus mykiss (rainbow trout)  
 C;Date: 13-Sep-1996 #sequence\_revision 13-Sep-1996 #text\_change 16-Jul-1999

GenCore version 5.1.3  
Copyright (c) 1993 - 2003 Compugen Ltd.

OM protein - protein search, using BW model

Run on: 4 February 13, 2003, 16:59:58 ; Search time 12 Seconds  
(without alignments)  
134,798 Million cell updates/sec

Title: US-09-756-690A-2

Perfect score: 209

Sequence: 1 HGGTGFITSDLSKQMBEAVRLFIEWLNGGDPSSGAPPPS 39

Scoring table: BLOSUM62

Gapop 10.0 , Gapext 0.5

Searched: 112892 seqs, 4176328 residues

Total number of hits satisfying chosen parameters: 112892

Minimum DB seq length: 0

Maximum DB seq length: 2000000000

Post-processing: Minimum Match 0%  
Maximum Match 100%  
Listing first 45 summaries

Database : SwissProt\_40.0\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

#### SUMMARIES

Result No.	Query	Match	Length	DB	ID	Description	
1	EXE4_HELSU	87	1	EXE5_HELSU	P26349	heloderma s	
2	200	95.7	39	1	EXE5_HELSU	P20394	heloderma h
3	97	46.4	71	1	GLUC_ICTPU	P04093	ictalurus p
4	97	46.4	103	1	GLUC_PIANCA	P15438	rana catesbe
5	93	44.5	71	1	GLUC_PIANCA	P81880	piaractus m
6	93	44.5	121	1	GLUC_CARAU	P79695	carassius a
7	92	44.0	30	1	GLUC_ANGUILA	P14521	anguilla an
8	89	42.6	78	1	GLUC_LEPSP	P09566	lepisosteus
9	88	42.1	151	1	GLUC_CHICK	P012277	gallus galli
10	87	41.6	29	1	GLUC_SECYCA	P09687	scylliorhinus
11	87	41.6	158	1	GLUC_PIG	P012474	sus scrofa
12	87	41.6	180	1	GLUC_BOVIN	P0122	bos tauris
13	87	41.6	180	1	GLUC_CAVPO	P05110	cavia porce
14	87	41.6	180	1	GLUC_HUMAN	P01215	homo sapien
15	87	41.6	180	1	GLUC_MASAU	P01235	mesocichetus
16	87	41.6	180	1	GLUC_MOUSE	P55095	mus musculus
17	87	41.6	180	1	GLUC_OCTDE	P22890	octodon deg
18	87	41.6	180	1	GLUC_RAT	P06833	rattus norvegicus
19	86	41.1	122	1	GLUC_LOPAM	P04042	lophius aeneus
20	84	40.2	29	1	GLUC_PLATEY	P23012	platichthys
21	84	40.2	29	1	GLUC_TORMA	P09567	torpedo marmoratus
22	84	40.2	96	1	GLUC_MYOSIC	P09686	myoxocephalus
23	83	39.7	36	1	GLU1_ORBNI	P81016	oreochromis
24	83	39.7	68	1	GLUC_ONCII	P07449	oncorhynchus
25	81	38.8	29	1	GLUC_CHIBR	P31227	chinchilla
26	81	38.8	124	1	GLUC_LOPAM	P18178	lophius aeneus
27	80	38.3	29	1	GLUC_RABIT	P25449	oryctolagus
28	80	38.3	69	1	GLUC_TORMA	P29794	canis familiaris
29	79	37.8	33	1	GLU2_ORBNI	P81027	oreochromis
30	75	35.9	29	1	GLUC_CALMI	P13139	callopterus
31	74	35.9	29	1	GLUC_DIDMA	P18108	didelphis marsupialis
32	74	35.4	29	1	GLUC_ANAPL	P01226	anas platyrhynchos
33	74	35.4	75	1	GLUC_AMICA	P33528	amia calva

RESULT 1						
EXP4_HELSU	HBLSU	STANDARD	PRT;	87 AA.		
1D_EXB4_HBLSU						
AC P26349						
DT 01-MAY-1992 (Rel. 22, Created)						
DT 15-JUL-1998 (Rel. 36, Last sequence update)						
DT 15-JUN-2002 (Rel. 41, Last annotation update)						
DE Exendin-4 precursor.						
OS Heloderma suspectum (Gila monster); Eukaryota; Chordata; Craniata; Vertebrata; Buteleostomi; Lepidosauria; Squamata; Scleroglossa; Anguimorpha; Helodermatidae; Heloderma						
OC						
OC						
NCBI_TaxID=8554;						
OX						
RN [1]						
SEQUENCE FROM N.A.						
RX MEDLINE#=97172477; PubMed=9020121;						
RA Chen Y.B., Drucker D.J.;						
RT "Tissue-specific expression of unique mRNAs that encode proglucagon-derived peptides or exendin 4 in the lizard." J. Biol. Chem. 272:4108-4115(1997).						
RN [2]						
SEQUENCE OF 48-86.						
RC TISSUE=Venom;						
RX MEDLINE#=92218391; PubMed=1313797;						
RA Eng J., Kleiman W.A., Singh L., Singh G., Raufman J.-P.;						
RT Isolation and characterization of exendin-4, an exendin-3 analogue, from Heloderma suspectum venom. Further evidence for an exendin receptor on dispersed acini from guinea pig pancreas." J. Biol. Chem. 267:7402-7405(1992).						
CC 1- FUNCTION: HAS A VIP SECRETIN-LIKE BIOLOGICAL ACTIVITY. INTERACTS WITH THE EXENDIN RECEPTOR.						
CC 1- SUBCELLULAR LOCATION: Secreted.						
CC 1- TISSUE SPECIFICITY: BELONGS TO THE GLUCAGON FAMILY.						
CC 1- SIMILARITY: BELOWS TO THE GLUCAGON FAMILY.						
-----						
This SWISS-PROT entry is copyright. It is produced through a collaboration between the Swiss Institute of Bioinformatics and the EMBL outstation - the European Bioinformatrics Institute. There are no restrictions on its use by non-profit institutions as long as its content is in no way modified and this statement is not removed. Usage by and for commercial entities requires a license agreement (See <a href="http://www.isb-sib.ch/announce/">http://www.isb-sib.ch/announce/</a> or send an email to license@isb-sib.ch).						
CC EMBL; U77113; AAB51130.1; -.						
DR InterPro: IP000032; Glucagon.						
DR Pfam: PF00123; hormone2; 1.						
SMART; SM0010; GLUCA; 1.						
PROSITE; PS02620; Glucagon; 1.						
KW Glucagon family; Toxin; Amidation; Signal.						
CC SIGNAL: 1 23 POTENTIAL.						
DR A42486; HWG4G.						
DR InterPro: IP000032; Glucagon.						
DR Pfam: PF00123; hormone2; 1.						
SMART; SM0010; GLUCA; 1.						
PROSITE; PS02620; Glucagon; 1.						
KW Glucagon family; Toxin; Amidation; Signal.						
CC SIGNAL: 1 23 POTENTIAL.						
DR A42486; HWG4G.						
DR InterPro: IP000032; Glucagon.						
DR Pfam: PF00123; hormone2; 1.						
SMART; SM0010; GLUCA; 1.						
PROSITE; PS02620; Glucagon; 1.						
KW Glucagon family; Toxin; Amidation; Signal.						
CC SIGNAL: 1 23 POTENTIAL.						
DR A42486; HWG4G.						
DR InterPro: IP000032; Glucagon.						
DR Pfam: PF00123; hormone2; 1.						
SMART; SM0010; GLUCA; 1.						
PROSITE; PS02620; Glucagon; 1.						
KW Glucagon family; Toxin; Amidation; Signal.						
CC SIGNAL: 1 23 POTENTIAL.						
DR A42486; HWG4G.						
DR InterPro: IP000032; Glucagon.						
DR Pfam: PF00123; hormone2; 1.						
SMART; SM0010; GLUCA; 1.						
PROSITE; PS02620; Glucagon; 1.						
KW Glucagon family; Toxin; Amidation; Signal.						
CC SIGNAL: 1 23 POTENTIAL.						
DR A42486; HWG4G.						
DR InterPro: IP000032; Glucagon.						
DR Pfam: PF00123; hormone2; 1.						
SMART; SM0010; GLUCA; 1.						
PROSITE; PS02620; Glucagon; 1.						
KW Glucagon family; Toxin; Amidation; Signal.						
CC SIGNAL: 1 23 POTENTIAL.						
DR A42486; HWG4G.						
DR InterPro: IP000032; Glucagon.						
DR Pfam: PF00123; hormone2; 1.						
SMART; SM0010; GLUCA; 1.						
PROSITE; PS02620; Glucagon; 1.						
KW Glucagon family; Toxin; Amidation; Signal.						
CC SIGNAL: 1 23 POTENTIAL.						
DR A42486; HWG4G.						
DR InterPro: IP000032; Glucagon.						
DR Pfam: PF00123; hormone2; 1.						
SMART; SM0010; GLUCA; 1.						
PROSITE; PS02620; Glucagon; 1.						
KW Glucagon family; Toxin; Amidation; Signal.						
CC SIGNAL: 1 23 POTENTIAL.						
DR A42486; HWG4G.						
DR InterPro: IP000032; Glucagon.						
DR Pfam: PF00123; hormone2; 1.						
SMART; SM0010; GLUCA; 1.						
PROSITE; PS02620; Glucagon; 1.						
KW Glucagon family; Toxin; Amidation; Signal.						
CC SIGNAL: 1 23 POTENTIAL.						
DR A42486; HWG4G.						
DR InterPro: IP000032; Glucagon.						
DR Pfam: PF00123; hormone2; 1.						
SMART; SM0010; GLUCA; 1.						
PROSITE; PS02620; Glucagon; 1.						
KW Glucagon family; Toxin; Amidation; Signal.						
CC SIGNAL: 1 23 POTENTIAL.						
DR A42486; HWG4G.						
DR InterPro: IP000032; Glucagon.						
DR Pfam: PF00123; hormone2; 1.						
SMART; SM0010; GLUCA; 1.						
PROSITE; PS02620; Glucagon; 1.						
KW Glucagon family; Toxin; Amidation; Signal.						
CC SIGNAL: 1 23 POTENTIAL.						
DR A42486; HWG4G.						
DR InterPro: IP000032; Glucagon.						
DR Pfam: PF00123; hormone2; 1.						
SMART; SM0010; GLUCA; 1.						
PROSITE; PS02620; Glucagon; 1.						
KW Glucagon family; Toxin; Amidation; Signal.						
CC SIGNAL: 1 23 POTENTIAL.						
DR A42486; HWG4G.						
DR InterPro: IP000032; Glucagon.						
DR Pfam: PF00123; hormone2; 1.						
SMART; SM0010; GLUCA; 1.						
PROSITE; PS02620; Glucagon; 1.						
KW Glucagon family; Toxin; Amidation; Signal.						
CC SIGNAL: 1 23 POTENTIAL.						
DR A42486; HWG4G.						
DR InterPro: IP000032; Glucagon.						
DR Pfam: PF00123; hormone2; 1.						
SMART; SM0010; GLUCA; 1.						
PROSITE; PS02620; Glucagon; 1.						
KW Glucagon family; Toxin; Amidation; Signal.						
CC SIGNAL: 1 23 POTENTIAL.						
DR A42486; HWG4G.						
DR InterPro: IP000032; Glucagon.						
DR Pfam: PF00123; hormone2; 1.						
SMART; SM0010; GLUCA; 1.						
PROSITE; PS02620; Glucagon; 1.						
KW Glucagon family; Toxin; Amidation; Signal.						
CC SIGNAL: 1 23 POTENTIAL.						
DR A42486; HWG4G.						
DR InterPro: IP000032; Glucagon.						
DR Pfam: PF00123; hormone2; 1.						
SMART; SM0010; GLUCA; 1.						
PROSITE; PS02620; Glucagon; 1.						
KW Glucagon family; Toxin; Amidation; Signal.						
CC SIGNAL: 1 23 POTENTIAL.						
DR A42486; HWG4G.						
DR InterPro: IP000032; Glucagon.						
DR Pfam: PF00123; hormone2; 1.						
SMART; SM0010; GLUCA; 1.						
PROSITE; PS02620; Glucagon; 1.						
KW Glucagon family; Toxin; Amidation; Signal.						
CC SIGNAL: 1 23 POTENTIAL.						
DR A42486; HWG4G.						
DR InterPro: IP000032; Glucagon.						
DR Pfam: PF00123; hormone2; 1.						
SMART; SM0010; GLUCA; 1.						
PROSITE; PS02620; Glucagon; 1.						
KW Glucagon family; Toxin; Amidation; Signal.						
CC SIGNAL: 1 23 POTENTIAL.						
DR A42486; HWG4G.					</td	

Best Local Similarity 100.0%; Pred. No. 8 1e-20; Matches 39; Conservative 0; Mismatches 0; Indels 0; Gaps 0; RN

Qy 1 HEGGTFTSDLSKQMEEAVLTFEVLKNGGPGSGAPPS 39  
Db 48 HEGGTFTSDLSKQMEEAVLTFEVLKNGGPGSGAPPS 86

RESULT 2

EXE3\_HELI0 ID HELHO STANDARD; PRT; 39 AA.  
AC P20394; DT 01-FEB-1991 (Rel. 17, Created)  
DT 01-FEB-1991 (Rel. 17, Last sequence update)  
DT 15-JUN-2002 (Rel. 41, Last annotation update)  
DE Exendin-3.  
OS Heloderma horridum horridum (Mexican beaded lizard).  
OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
Lepidosaurs; Squamata; Scleroglossa; Anguimorpha; Helodermatidae;  
Heloderma.  
NCBI\_TaxID=8552; RN [1]

RP SEQUENCE.  
RC TISSUE="Venom"; MEDLINE=91056067; PubMed=1700785;  
RX Eng J.; Andrew P.C., Kleiman W.A., Singh L., Raufman J.-P.; RA  
RT "Purification and structure of exendin-3, a new pancreatic  
secretagogue isolated from Heloderma horridum venom.",  
RL J. Biol. Chem. 265:20259-20262 (1990).  
CC -; FUNCTION: HAS A VIP/SECRETIN-LIKE BIOLOGICAL ACTIVITY. INTERACTS  
WITH THE EXENDIN RECEPTOR.  
CC -; SUBCELLULAR LOCATION: Secreted.  
CC -; TISSUE SPECIFICITY: Produced by the venomous gland.  
DR PIR: A23674; HNC3H32.  
HSSP; P01275; 1BH0  
InterPro; IPR00532; Glucagon.  
Pfam; PF00123; hormone2; 1.  
SMART; SM00070; SM00070; GLUCA; 1.  
PROSITE; PS00260; GLUCA; 1.  
RN Glucagon family; Toxin; Amidation.  
MOD RES 39 39 AMIDATION.  
SEQUENCE 39 AA; A42451D3A4B1D1B9 CRC64;

Query Match 95.7%; Score 200; DB 1; Length 39;  
Best Local Similarity 94.9%; Pred. No. 4.5e-19;  
Matches 37; Conservative 1; Mismatches 1; Indels 0; Gaps 0; RN

Qy 1 HEGGTFTSDLSKQMEEAVLTFEVLKNGGPGSGAPPS 39  
Db 1 HSDGTFTSDLSKQMEEAVLTFEVLKNGGPGSGAPPS 39

RESULT 3

GLUC\_ICIPU ID GLUC\_ICIPU STANDARD; PRT; 71 AA.  
AC P01093; DT 01-NOV-1986 (Rel. 03, Created)  
DT 01-MAR-1989 (Rel. 10, Last sequence update)  
DE Glucagon precursor (Fragment).  
OS Ictalurus punctatus (Channel catfish).  
Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
Actinopterygi; Neopterygi; Teleostei; Ostariophys; Siluriformes;  
Ictaluridae; Ictalurus.  
NCBI\_TaxID=1998; RN [1]

RP SEQUENCE.  
RC TISSUE="Pancreas"; MEDLINE=87156787; PubMed=31030321;  
RX Hoesein N.M., Mahrenholz A.M., Andrews P.C., Gurd R.S.;  
RT "Biological activities of catfish glucagon and glucagon-like  
peptide.",

Best Local Similarity 100.0%; Pred. No. 8 1e-20; Matches 39; Conservative 0; Mismatches 0; Indels 0; Gaps 0; RN  
[2]  
Biochem. Biophys. Res. Commun. 143:87-92 (1987).  
RN SEQUENCE.  
RC TISSUE="Pancreas"; MEDLINE=85157516; PubMed=3830546;  
RX Andrews P.C., Ronner P.; RR "Isolation and structures of glucagon and glucagon-like peptide from catfish pancreas"; J. Biol. Chem. 260:3910-1914 (1985).  
CC -; FUNCTION: PROMOTES HYDROLYSIS OF GLYCOGEN AND LIPIDS, AND RAISES THE BLOOD SUGAR LEVEL.  
CC -; INDUCTION: PRODUCED IN THE A CELLS OF THE ISLETS OF LANGERHANS  
IN RESPONSE TO A DROP IN BLOOD SUGAR CONCENTRATION.  
CC -; MISCELLANEOUS: X'S IN THE SEQUENCE WERE INCLUDED BY HOMOLOGY WITH AMERICAN GOOSEFISH SEQUENCES.  
CC -; SIMILARITY: BELONGS TO THE GLUCAGON FAMILY.  
DR PIR; A05166; QCIDC.  
HSSP; P01274; 1GCR.  
InterPro; IPR000532; Glucagon.  
Pfam; PF00123; hormone2; 2.  
SMART; SM00070; GLUCA; 2.  
DR PROSITE; PS00260; GLUCA; 2.  
RN Glucagon family; Hormone.  
FT NON\_TER 1 1 GLUCAGON.  
FT PEPTIDE 1 29 GLUCAGON-LIKE PEPTIDE.  
FT PEPTIDE 38 71 GLUCAGON.  
FT CONFLICT 53 53 E -> D (IN REF. 2).  
FT NON\_TER 71 71 GLUCAGON.  
SQ SEQUENCE 71 AA; 8173 MW; 246887E79AD981A8F CRC44;  
Query Match 46.4%; Score 97; DB 1; Length 71;  
Best Local Similarity 51.6%; Pred. No. 1e-05;  
Matches 16; Conservative 8; Mismatches 7; Indels 0; Gaps 0; RN  
[1]  
RESULT 4  
GLUC\_RANCA ID GLUC\_RANCA STANDARD; PRT; 103 AA.  
AC P1541B; P15439; P15440;  
DT 01-APR-1990 (Rel. 14, Created)  
DT 01-JUL-1993 (Rel. 26, Last sequence update)  
DT 01-JUL-1993 (Rel. 26, Last annotation update)  
DR Glucagon Precursor (Fragment).  
Rana catesbeiana (Bull frog).  
OS Amphibia; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
OC Bokaryota; Batrachia; Anura; Neobatrachia; Ranidae; Ranidae; Rana.  
NCBI\_TaxID=8400; RN [1]  
RP SEQUENCE.  
RC TISSUE="Pancreas"; MEDLINE=88257102; PubMed=3260236;  
RX Pollock H.G., Hamilton J.W., Rouse J.B., Ebner K.B., Rawitch A.B.; RR "Isolation of Peptide hormones from the pancreas of the bullfrog (Rana catesbeiana). Amino acid sequences of pancreatic polypeptide, oxyntomodulin, and two glucagon-like peptides."; RL J. Biol. Chem. 263:9746-9751 (1988).  
CC -; FUNCTION: PROMOTES HYDROLYSIS OF GLYCOGEN AND LIPIDS, AND RAISES THE BLOOD SUGAR LEVEL.  
CC -; INDUCTION: PRODUCED IN THE A CELLS OF THE ISLETS OF LANGERHANS  
IN RESPONSE TO A DROP IN BLOOD SUGAR CONCENTRATION.  
CC -; MISCELLANEOUS: X'S IN THE SEQUENCE WERE INCLUDED BY HOMOLOGY WITH OTHER SPECIES SEQUENCES.  
CC -; SIMILARITY: BELONGS TO THE GLUCAGON FAMILY.  
DR PIR; B20891; QCGB.  
HSSP; P01274; 1GCR.  
InterPro; IPR000532; Glucagon.  
PRINTS; PR00275; GLUCAGON.  
SMART; SM00070; GLUCA; 3.  
DR PROSITE; PS00260; GLUCA; 3.  
RN [1]

RESULT 5

Query Match Score 97; DB 1; length 103;  
Best Local Similarity 51.6%; Pred. No. 1.5e-05;  
Matches 16; Conservative 7; Mismatches 8; Indels 0; Gaps 0;

Qy 1 HGGTFTSDLSKQMBEAVRLFIEWLNGGP 31  
Db 39 HADGTTFTSDMSYLEEKAKEFVDWLIKGRP 69

RESULT 5

GLUC\_PIAME STANDARD; PRT; 71 AA.

ID\_GLUC\_PIAME STANDARD; PRT; 71 AA.

AC P81880; 39. Created

DT 30-MAY-2000 (Rel. 39. Last sequence update)

DT 30-MAY-2000 (Rel. 39. Last annotation update)

DT 30-MAY-2000 (Rel. 39. Last annotation update)

DE Glucagon precursor (Fragment).

OS Piaractus mesopotamicus (Pacu).

OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Buteleostomi; Actinopterygii; Neopterygii; Teleostei; Ostariophysii; Characiformes; Characidae; Piaractidae; Piaractus.

NCBI\_TaxID=42528; [1]

RN SEQUENCE.

RP TISSUE="Pancreas"; PubMed=10327603;

RX MEDLINE=99229587; PubMed=10327603;

RA de Lima J.A., Oliveira B., Conlon J.M.;

RT "Purification and characterization of insulin and peptides derived from proglucagon and prosomatosatin from the fruit-eating fish, the pacu Piaractus mesopotamicus";

RL Comp. Biochem. Physiol. 128B:127-135 (1995).

CC -1- FUNCTION: PROMOTES HYDROLYSIS OF GLYCOCEN AND LIPIDS, AND RAISES THE BLOOD SUGAR LEVEL.

CC -1- INDUCTION: PRODUCED IN THE A CELLS OF THE ISLETS OF LANGERHANS IN RESPONSE TO A DROP IN BLOOD SUGAR CONCENTRATION.

CC -1- MISCELLANEOUS: X'S IN THE SEQUENCE WERE INCLUDED BY HOMOLOGY WITH OTHER FISH SEQUENCES.

CC -1- SIMILARITY: BELONGS TO THE GLUCAGON FAMILY.

DR HSSP; P01274; IGCN.

DR InterPro; IPR00532; Glucagon.

DR Pfam; PF00123; hormone2; 2.

DR PRINTS; PRO0075; GLUCAGON.

DR SMART; SM00070; GLUCAG.

DR PROSITE; PS00266; GLUCAG; 2.

KW Glucagon family; Hormone.

PT NON\_TER 1 1

PT PEPTIDE 1 29 GLUCAGON.

PT PEPTIDE 38 71 GLUCAGON-LIKE PEPTIDE.

PT NON\_TER 71 71 F6A3C2ADD980655 CRC64;

SQ SEQUENCE 71 AA; 8146 MW; F6A3C2ADD980655 CRC64;

Query Match Score 93; DB 1; length 71;  
Best Local Similarity 48.4%; Pred. No. 3.2e-05;  
Matches 15; Conservative 9; Mismatches 7; Indels 0; Gaps 0;

Qy 1 HGGTFTSDLSKQMBEAVRLFIEWLNGGP 31  
Db 38 HADGTTFTSDMSYLEEKAKEFVDWLIKGRP 69

RESULT 6

GLUC\_CARAU STANDARD; PRT; 121 AA.

ID\_GLUC\_CARAU STANDARD; PRT; 121 AA.

AC P78655; [1]

DT 01-NOV-1997 (Rel. 35. Last sequence update)

DE Glucagon precursor [Contains: Glicentin-related polypeptide (GRPP); Carassius auratus (Goldfish); Carassius auratus (Goldfish); Metazoa; Chordata; Craniata; Vertebrata; Buteleostomi; Actinopterygii; Neopterygii; Teleostei; Ostariophysii; Cypriniformes; Cyprinidae; Carassius; NCBI\_TaxID=7957; [1]

DP 01-NOV-1997 (Rel. 35. Last sequence update)

DT 16-OCT-2001 (Rel. 40. Last annotation update)

DE Glucagon precursor [Contains: Glicentin-related polypeptide (GRPP); Carassius auratus (Goldfish);

OS Carassius auratus (Goldfish).

OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Buteleostomi; Actinopterygii; Neopterygii; Teleostei; Ostariophysii; Cypriniformes; Cyprinidae; Carassius.

CC SEQUENCE FROM N.A.

RA Yuen T.T.H., Mok P.Y., Chow B.K.C.; GenBank/GenBank/DBJ databases.

CC Submitted (FEB-1997) to the EMBL/GenBank/DBJ databases.

CC -!- FUNCTION: PROMOTES HYDROLYSIS OF GLYCOCEN AND LIPIDS, AND RAISES THE BLOOD SUGAR LEVEL.

CC -!- SIMILARITY: BELONGS TO THE GLUCAGON FAMILY.

CC -!- SIMILARITY: BELONGS TO THE GLUCAGON FAMILY.

CC This SWISS-PROT entry is copyright. It is produced through a collaboration between the Swiss Institute of Bioinformatics and the EMBL outstation - the European Bioinformatics Institute. There are no restrictions on its use by non-profit institutions as long as its content is in no way modified and this statement is not removed. Usage by and for commercial entities requires a license agreement. (See <http://www.isb-sib.ch/announce/> or send an email to license@isb-sib.ch).

CC EMBL; U65528; AAB9563.1; -.

CC HSSP; P01274; IGCN.

CC InterPro; IPR00532; Glucagon.

CC Pfam; PF00123; hormone2; 2.

CC PRINTS; PRO0275; GLUCAGON.

CC SMART; SM00070; GLUCAG; 2.

CC PROSITE; PS00360; GLUCAGON.

CC KW Glucagon family; Hormone; Cleavage on pair of basic residues; Signal.

CC SIGNAL 1 21 POTENTIAL POLYPEPTIDE.

CC FT PEPTIDE 22 47 GLUCENTIN-RELATED POLYPEPTIDE.

CC FT PEPTIDE 50 78 GLUCAGON.

CC FT PROPER 90 95 GLUCAGON.

CC FT PEPTIDE 88 121 GLUCAGON-LIKE PEPTIDE.

CC SQ SEQUENCE 121 AA; 13527 MW; 5C1D4B2C1D26B9C6 CRC64;

CC Query Match Score 93; DB 1; length 121;  
CC Best Local Similarity 48.4%; Pred. No. 5.8e-05;  
CC Matches 15; Conservative 8; Mismatches 8; Indels 0; Gaps 0;

Qy 1 HGGTFTSDLSKQMBEAVRLFIEWLNGGP 31  
Db 88 HADGTTFTSDMSYLEEKAKEFVDWLIKGRP 69

RESULT 7

GLUC\_ANGAN STANDARD; PRT; 30 AA.

ID\_GLUC\_ANGAN STANDARD; PRT; 30 AA.

AC P41521; [1]

DT 01-NOV-1995 (Rel. 32. Created)

DT 01-NOV-1995 (Rel. 32. Last sequence update)

DT 01-NOV-1995 (Rel. 32. Last annotation update)

DT 01-NOV-1995 (Rel. 32. Last annotation update)

DR Glucagon-like Peptide (GLP).

OS Anguilla anguilla (European eel).

OS Anguilla rostrata (American eel).

OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Buteleostomi; Actinopterygii; Neopterygii; Teleostei; Anguillidae; Anguilla.

NCBI\_TaxID=7936; 7938; [1]

RP SEQUENCE.

RX TISSUE=Pancreas;

RX MEDLINE=91340068; PubMed=1874185;

RA Conlon J.M., Andrews P.C., Thim L., Moon T.W.;

RR The primary structure of Glucagon-like peptide but not insulin has been conserved between the American eel, Anguilla rostrata and the European eel, Anguilla anguilla.";

RR Gen. Comp. Endocrinol. 82:23-32 (1991).

CC -!- SIMILARITY: BELONGS TO THE GLUCAGON FAMILY.

DR PIR; B61125; B61125.  
 DR PIR; C61125; C61125.  
 DR HSSP; P01275; 1BH0.  
 DR InterPro; IPR000532; Glucagon.  
 Pfam; PF00123; hormone2; 1.  
 SMART; SM00070; GLUCAGON.  
 PROSITE; PS00260; GLUCAGON; 1.  
 KW Glucagon family; Amidation.  
 MOD RES 30 30 AMIDATION.  
 PT SEQUENCE 30 AA: 3376 MW: 592DA5EABD6E49D0 CRC64;  
 Query Match 44.0%; Score 92; DB 1; Length 30;  
 Best Local Similarity 48.3%; Pred. No. 1.6e-05;  
 Matches 14; Conservative 8; Mismatches 7; Indels 0; Gaps 0;  
 Qy 1 HEGGTFTSDLSKQMEBAVRLPEWIKNG 29  
 Db 1 HAEGTFTSDVSYLQDQAKEYFWSWLKTG 29

RESULT 8  
 GLUC\_LBSPP\_LBSP STANDARD; PRT; 78 AA.  
 ID GLUC\_LBSP  
 AC P09766;  
 DT 01-MAR-1989 (Rel. 10, Created)  
 DT 01-NOV-1990 (Rel. 16, Last sequence update)  
 DT 16-OT-2001 (Rel. 40, Last annotation update)  
 DB Glucagon Precursor [Contains: Glucagon; Glucagon-36 (Oxyntomodulin); Glucagon-like Peptide (Fragment).  
 OS Lepisosteus platulus (Alligator gar) (Atractosteus spatula).  
 OC Bokaryota; Metazoa; Chordata; craniata; Vertebrata; Euteleostomi; Actinopterygii; Neopterygii; Semionotiformes; Lepisosteidae; Lepisosteus.  
 OX NCBI\_TAXID=917;  
 RN 1  
 RP SEQUENCE OF 1-36 AND 45-78.  
 TISSUE=Pancreas;  
 MEDLINE=198796; PubMed=33282974;  
 RA Pollock H.G., Kimmel J.R., Ebner K.B., Hamilton J.W., Rouse J.B., Lance V., Rawitch A.B.;  
 RT "Isolation of alligator gar (Lepisosteus spatula) glucagon, oxyntomodulin, and glucagon-like peptide; amino acid sequences of oxyntomodulin and glucagon-like peptide.",  
 RT Gen. Comp. Endocrinol. 69:133-140(1988).  
 RN [2]  
 RP PRELIMINARY SEQUENCE OF 1-23.  
 TISSUE=Pancreas;  
 MEDLINE=88030591; PubMed=33111873;  
 RA Pollock H.G., Kimmel J.R., Hamilton J.W., Rouse J.B., Ebner K.B., Lance V., Rawitch A.B.;  
 RT "Isolation and structure of alligator gar (Lepisosteus spatula) insulin and pancreatic polypeptide";  
 RL Gen. Comp. Endocrinol. 97:375-382(1997).  
 CC -!- FUNCTION: PROMOTES HYDROLYSIS OF GLYCOCEN AND LIPIDS, AND RAISES THE BLOOD SUGAR LEVEL.  
 CC -!- INDUCTION: PRODUCED IN THE A CELLS OF THE ISLETS OF LANGERHANS IN RESPONSE TO A DROP IN BLOOD SUGAR CONCENTRATION.  
 CC -!- MISCELLANEOUS: X'S IN THE SEQUENCE WERE INCLUDED BY HOMOLOGY WITH AMERICAN GOLDFISH SEQUENCES.  
 CC -!- SIMILARITY: BELONGS TO THE GLUCAGON FAMILY.  
 DR PIR; S06339; GCGX.  
 DR InterPro; IPR000532; Glucagon.  
 Pfam; PF00123; hormone2; 2.  
 DR HSSP; P01274; IGCN.  
 KW Glucagon family; Hormone.  
 PT PEPTIDE 1 29 GLUCAGON-36.  
 PT PEPTIDE 1 36 GLUCAGON-LIKE PEPTIDE.  
 PT PEPTIDE 45 78 GLUCAGON-LIKE PEPTIDE.  
 SEQUENCE 78 AA: 8990 MW: 301064927159450 CRC64;

Query Match 42.6%; Score 89; DB 1; Length 78;  
 Best Local Similarity 44.4%; Pred. No. 0.00011; Gaps 0;  
 Matches 13; Conservative 9; Mismatches 7; Indels 0; Gaps 0;  
 Qy 1 HEGGTFTSDLSKQMEBAVRLPEWIKNG 29  
 Db 45 HADGTYSVDSVSYLQDQAKEYFWSWLKG 73

RESULT 9  
 GLUC\_CHICK STANDARD; PRT; 151 AA.  
 ID GLUC\_CHICK  
 AC P01277;  
 DT 21-JUL-1986 (Rel. 01, Created)  
 DT 01-AUG-1990 (Rel. 15, Last sequence update)  
 DT 15-JUL-1999 (Rel. 38, Last annotation update)  
 DB Glucagon Precursor.  
 OS Gallus gallus (Chicken). and Meleagris gallopavo (Common turkey).  
 OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Butelostomi; Archosauria; Aves; Neognathae; Galliformes; Phasianidae; Gallus.  
 OC NCBI\_TAXID=9031; 9103;  
 OX RN [1]  
 RP SEQUENCE FROM N.A.  
 RC SPECIES=Chicken; TISSUE=Pancreas;  
 MEDLINE=90243942; PubMed=2330135;  
 RX RA Hasegawa S., Terazono K., Nata K., Takada T., Yamamoto H., Okamoto H.;  
 PT "Nucleotide sequence determination of chicken glucagon precursor DNA. Chicken proglucagon does not contain glucagon-like peptide III.";  
 RT FEBS Lett. 264:117-120(1990).  
 RN [2]  
 RP SEQUENCE OF 55-83.  
 RC SPECIES=Chicken;  
 MEDLINE=16062271; PubMed=1194290;  
 RX RA Pollock H.G., Kimmel J.R.,  
 RT "Chicken glucagon. Isolation and amino acid sequence studies.",  
 RL J. Biol. Chem. 250:9377-9380(1975).  
 RN [3]  
 RP COMPOSITION, AND SEQUENCE OF 55-83.  
 RC SPECIES=M.Gallopavo;  
 MEDLINE=73074118; PubMed=4645932;  
 RX RA Markussen J., Frandsen E.K., Heding L.G., Sundby F.;  
 RT "Turkey glucagon: crystallization, amino acid composition and immunology.",  
 RL Horm. Metab. Res. 4:360-363 (1972).  
 CC -!- FUNCTION: PROMOTES HYDROLYSIS OF GLYCOCEN AND LIPIDS, AND RAISES THE BLOOD SUGAR LEVEL.  
 CC -!- INDUCTION: PRODUCED IN THE A CELLS OF THE ISLETS OF LANGERHANS IN RESPONSE TO A DROP IN BLOOD SUGAR CONCENTRATION.  
 CC -!- MISCELLANEOUS: THE COMPOSITION OF TURKEY GLUCAGON APPEARS TO BE IDENTICAL WITH CHICKEN. CHICKEN PREPROGLUCAGON DOES NOT CONTAIN GLUCAGON-LIKE PEPTIDE II.  
 CC -!- SIMILARITY: BELONGS TO THE GLUCAGON FAMILY.  
 CC This SWISS-PROT entry is copyright. It is produced through a collaboration between the Swiss Institute of Bioinformatics and the EMBL outstation - the European Bioinformatics Institute. There are no restrictions on its use by non-profit institutions as long as its content is in no way modified and this statement is not removed. Usage by and for commercial entities requires a license agreement. (See http://www.isb-sib.ch/announce/ or send an email to license@isb-sib.ch).  
 CC EMBL; Y07539; CAA68827; 1; -.  
 DR PIR; S09992; GCGC.  
 DR PIR; A91740; A91740.  
 DR HSSP; P01274; IGCN.  
 DR InterPro; IPR0000532; Glucagon.  
 DR Pfam; PF00123; hormone2; 2.  
 DR PRINTS; PR00275; GLUCAGON.

DR SMART; SM00070; GLUCA; 2.  
 DR PROSTIE; PS00260; GLUCAGON; 3.  
 KW Glucagon family; Hormone; Cleavage on pair of basic residues; Signal;  
 Amidation.  
 FT SIGNAL 1 22  
 FT CHAIN 23 151 PROGLUCAGON.  
 FT PEPTIDE 55 83 GLUCAGON.  
 FT PROTEP 86 118 GLUCAGON-LIKE PEPTIDE.  
 FT PEPTIDE 118 147 GLUCAGON-LIKE PEPTIDE.  
 FT MOD\_RES 147 147 AMIDATION (G-148 PROVIDE AMIDE GROUP).  
 SQ SEQUENCE 151 AA; 17520 MW; BECD753COAB5 CRC64;

Query Match 42.1%; Score 88; DB 1; Length 151;  
 Best Local Similarity 51.7%; Pred. No. 0.00032; Indels 0; Gaps 0;  
 Matches 15; Conservative 6; Mismatches 8; Delins 0; Gaps 0;

Qy 1 HGEQTFTSDLSKQMBEAVRLFIWLNKG 29  
 Db 118 HAEGTTSIDTSTYLEGQAQAKFIAWLNG 146

---

RESULT 10  
 ID GLUC\_SCYCA STANDARD; PRT; 29 AA.  
 AC P09687;  
 DT 01-MAR-1989 (Rel. 10. Created)  
 DT 01-MAR-1989 (Rel. 10. Last sequence update)  
 DT 01-JAN-1990 (Rel. 13. Last annotation update)  
 DB Glucagon.  
 OS Scyliorhinus canicula (Spotted dogfish) (Spotted catshark)  
 OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Chondrichthyes;  
 OC Elasmobranchii; Galeomorphii; Galeoidea; Carcharhiniformes;  
 OC Scyliorhinidae; Scyliorhinus;  
 OX NCBI; TAXID=7830;  
 RN [1]  
 RP SEQUENCE.  
 RC TISSUE=Pancreas;  
 RX MEDLINE=87119051; PubMed=3560517;  
 RA Conlon J.M.; O'Toole L.; Thim L.;  
 RT "Primary structure of glucagon from the gut of the common dogfish  
 (Scyliorhinus canicula)."  
 RL FEBS Lett. 214:50-55 (1987).  
 CC -I- FUNCTION: PROMOTES HYDROLYSIS OF GLYCOGEN AND LIPIDS, AND RAISES  
 THE BLOOD SUGAR LEVEL.  
 CC -I- INDUCTION: PRODUCED IN THE A CELLS OF THE ISLETS OF LANGERHANS  
 IN RESPONSE TO A DROP IN BLOOD SUGAR CONCENTRATION.  
 CC -I- SIMILARITY: BELONGS TO THE GLUCAGON FAMILY.  
 DR PIR; A26992; GCF.  
 DR HSSP; P01274; 1GDN.  
 DR InterPro; IPR00532; Glucagon.  
 DR Pfam; PF00123; hormone2; 1.  
 DR PRINTS; PR00275; GLUCAGON.  
 DR SMART; SM00070; GLUCA; 1.  
 DR PROSTIE; PS00260; GLUCAGON; 1.  
 KW Glucagon family; Hormone;  
 SEQUENCE 29 AA; 3529 MW; 6FA96392086F0226 CRC64;

Query Match 41.6%; Score 87; DB 1; Length 29;  
 Best Local Similarity 53.6%; Pred. No. 6.7E-05; Indels 0; Gaps 0;  
 Matches 15; Conservative 4; Mismatches 9; Delins 0; Gaps 0;

Qy 1 HGEQTFTSDLSKQMBEAVRLFIWLNKG 28  
 Db 1 HSEGTFTSDLSKQMBEAVRLFIWLNKG 28

---

RESULT 11  
 ID GLUC\_PIG STANDARD; PRT;  
 AC P01274;  
 DT 21-JUL-1986 (Rel. 01. Created)  
 DT 01-NOV-1990 (Rel. 16. Last sequence update)  
 DT 16-OCT-2001 (Rel. 40. Last annotation update)

FT	HELIX	46	55					
FT	TURN	56	57					
SQ	SEQUENCE	158 AA;	18212 MW;	28CCFC257F333B2	CRC64;			
Query Match		41.6%	Score 87;	DB 1;	Length 158;			
Best Local Similarity		55.2%	Pred. No.	0.00045;				
Matches		16;	Conservative	4;	Mismatches	9;	Indels	0;
Qy	1	HGEGTFTSDLSKQMEBAEVLFIENLKG 29						
Db	18	HAEGTFTSDVSYLEGQAKETIAWLVKG 106						
RESULT 12								
GLUC BOVIN		STANDARD;	PRT;	180 AA.				
AC	P01272;							
DT	21-JUL-1986	(Rel. 01, Created)						
DT	13-AUG-1987	(Rel. 05, Last sequence update)						
DT	15-JUN-2002	(Rel. 41, Last annotation update)						
DE	Glucagon precursor [Contains: Glicentin-related polypeptide (GRPP); Glucagon; Glucagon-like peptide 1 (GLP1); Glucagon-like peptide 2 (GLP2)].							
GGC								
GN								
OS	Bos taurus (Bovine).							
OC	Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Buteleostomi; Mammalia; Eutheria; Cetartiodactyla; Ruminantia; Pecora; Bovidae; Bovidae; Bovinae; Bos.							
NCBI_TAXID	9913;							
RN								
RP	SEQUENCE FROM N.A.							
RX	Medline=83499946; PubMed=6577429;							
RX	Lopez L.C., Prazier M.L., Su C.-J., Kumar A., Saunders G.F.,							
RT	"Mammalian pancreatic preproglucagon contains three Glicentin-related peptides."							
PT								
PT	Proc. Natl. Acad. Sci. U.S.A. 80:5485-5489 (1983).							
RN								
RP	SEQUENCE FROM N.A.							
RX	Medline=71166445; PubMed=5102927;							
RX	Bromer W.W., Boucher M.B., Koffenberger J.B. Jr.;							
RT	"Amino acid sequence of bovine Glucagon.";							
RT	J. Biol. Chem. 246:2822-2827(1971).							
RN								
RP	STRUCTURE BY NMR OF 53-81.							
RX	Medline=71166445; PubMed=6631957;							
RX	Braun W., Wider G., Lee K.H., Wuthrich K.;							
RA	"Conformation of Glucagon in a lipid-water interphase by 1H nuclear magnetic resonance.";							
RT	J. Mol. Biol. 169:921-948 (1983).							
RT	"-1- FUNCTION: GLUCAGON PROMOTES HYDROLYSIS OF GLYCOGEN AND LIPIDS, AND RAISES THE BLOOD SUGAR LEVEL.							
CC	"-1- FUNCTION: GLP2 STIMULATES INTESTINAL GROWTH AND UPREGULATES VILLUS HEIGHT IN THE SMALL INTESTINE, CONCOMITANT WITH INCREASED CRYPT CELL PROLIFERATION AND DECREASED ENTEROCYTE APOPTOSIS.							
CC	"-1- INDUCTION: PRODUCED IN THE A CELLS OF THE ISLETS OF LANGERHANS IN RESPONSE TO A DROP IN BLOOD SUGAR CONCENTRATION.							
CC	"-1- SIMILARITY: BELONGS TO THE GLUCAGON FAMILY.							
CC	This SWISS-PROT entry is copyright. It is produced through a collaboration between the Swiss Institute of Bioinformatics and the EMBL outstation - the European Bioinformatics Institute. There are no restrictions on its use by non-profit institutions as long as its content is in no way modified and this statement is not removed. Usage by and for commercial entities requires a license agreement (See <a href="http://www.isb-sib.ch/announce/">http://www.isb-sib.ch/announce/</a> or send an email to license@isb-sib.ch).							
CC	EMBL: K00107; AAA30538; 1; -.							
CC	PIR: A01319; GBIO.							
CC	PDB: 1RXX; 13-PEB-02.							
CC	InterPro: IPR00532; Glucagon.							
CC	Pfam: PF00123; hormone2; 3.							
CC	PRINTS: PR00775; GLUCAGON.							
CC	SMART: SM00070; GLUCA_3.							
DR	DRB00014; BAA00010.1/ -.							
DR	PDB: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							
DR	DRPIR: A24856; GCGP.							
DR	DRP00123; hormone2; 3.							
DR	DRS00070; GLUCAGON.							



01-FEB-1996 (Rel. 33, Last sequence update)  
 DT 16-OCT-2001 (Rel. 40, Last annotation update)  
 DE Glucagon Precursor [Contains: Glucagon-related polypeptide (GGRP);  
 DE Glucagon; Glucagon-like peptide 1 (GLP1); Glucagon-like peptide 2  
 DE (GLP2)].  
 GCG  
 GN Mesocricetus auratus (Golden hamster)  
 OS Mammalia: Eutheria; Metacota; Chordata; Craniata; Vertebrata; Euteleostomi;  
 OC Mesocricetus auratus (Golden hamster)  
 OC Mammalia: Eutheria; Rodentia; Sciurognathi; Muridae; Cricetinae;  
 OC Mesocricetus.  
 RN [1]  
 RP SEQUENCE FROM N. A.  
 RX MEDLINE=83167563; PubMed=6835407;  
 RA Bell G.J., Santarre R.F., Mullerbach G.T.;  
 RT "Hamster preproglucagon contains the sequence of glucagon and two  
 related peptides";  
 RL Nature 302:716-718(1983).  
 RN [2]  
 RP REVISIONS TO 12-15.  
 RA Bell G.J.  
 RL Submitted (XXX 1985) to the EMBL/GenBank/DDBJ databases.  
 CC -I- FUNCTION: GLUCAGON PROMOTES HYDROLYSIS OF GLYCOGEN AND LIPIDS, AND  
 CC RAISES THE BLOOD SUGAR LEVEL.  
 CC -I- FUNCTION: GLP2 STIMULATES INTESTINAL GROWTH AND UPREGULATES VILLUS  
 CC HEIGHT IN THE SMALL INTESTINE, CONCOMITANT WITH INCREASED CRYPT  
 CC CELL PROLIFERATION AND DECREASED ENTEROCYTE APOPTOSIS.  
 CC -I- INDUCTION: PRODUCED IN THE A CELLS OF THE ISLETS OF LANGERHANS  
 CC IN RESPONSE TO A DROP IN BLOOD SUGAR CONCENTRATION.  
 CC -I- SIMILARITY: BELONGS TO THE GLUCAGON FAMILY.  
 CC -----  
 CC This SWISS-PROT entry is copyright. It is produced through a collaboration  
 CC between the Swiss Institute of Bioinformatics and the EMBL outstation -  
 CC the European Bioinformatics Institute. There are no restrictions on its  
 CC use by non-profit institutions as long as its content is in no way  
 CC modified and this statement is not removed. Usage by and for commercial  
 CC entities requires a license agreement (see <http://www.isb-sib.ch/announce/>  
 CC or send an email to license@isb-sib.ch).  
 CC -----  
 DR EMBL; J00059; AA37074.1; -  
 DR PIR; A01539; GCHY.  
 DR HSSP; P01274; IGEN.  
 DR InterPro; IPR000332; Glucagon.  
 DR Pfam; PF00123; Hormone2; 3.  
 DR PRINTS; PR00275; GLUCAGON.  
 DR SMART; SM00070; GLUCA; 3.  
 DR PROSITE; PS00260; GLUCAGON; 4.  
 KW Glucagon family; Hormone; Cleavage on pair of basic residues; Signal  
 FT SIGNAL; 1 20 GLICENTIN-RELATED POLYPEPTIDE.  
 FT PEPTIDE; 21 50 GLUCAGON.  
 FT PEPTIDE; 53 81 GLUCAGON-LIKE PEPTIDE 1.  
 FT PEPTIDE; 92 128 GLUCAGON-LIKE PEPTIDE 2.  
 FT PEPTIDE; 146 178 GLUCAGON-LIKE PEPTIDE 2.  
 SQ SEQUENCE 180 AA; 20954 MW; 02791149D7AADD4B CRC64;  
 Query Match 41.6%; Score 87; DB 1; Length 180;  
 Best Local Similarity 55.2%; Pred. No. 0.00052;  
 Matches 16; Conservative 4; Mismatches 9; Indels 0; Gaps 0;  
 Qy 1 HGEGTFTSDLSQMBEAVLPIWLRG 29  
 Db 98 HAECTFTSDVASSYLEGQAAKEFTAWLVRG 126

Search completed: February 13, 2003, 17:11:22  
 Job time : 30 SECs

Copyright (c) 1993 - 2003 Compugen Ltd.

OM protein - protein search, using sw model

Run on: February 13, 2003, 17:09:48 ; Search time 34 Seconds (without alignments)

Scoring table: BLOSUM62

Title: US-09-756-690A-2

Perfect score: 209

Sequence: 1 HCGTPTSDLQKMEAVRLFIEWLNGPSSGAPPS 39

Searched: 671580 seqs, 20604715 residues

Total number of hits satisfying chosen parameters: 671580

Post-processing: Minimum Match 0%

Database : SPREMBL 21;\*

Minimum DB seq length: 0

Maximum DB seq length: 2000000000

Post-processing: Minimum Match 0%

Maximum Match 100%

Listing first 45 summaries

Score Match Length DB ID

1: sp\_arbrea:\*

2: sp\_bacteria:\*

3: sp\_fungi:\*

4: sp\_human:\*

5: sp\_invertebrate:\*

6: sp\_mammal:\*

7: sp\_mhc:\*

8: sp\_organelle:\*

9: sp\_phage:\*

10: sp\_plant:\*

11: sp Rodent:\*

12: sp\_virus:\*

13: sp\_vertebrate:\*

14: sp\_unclassified:\*

15: sp\_xvirus:\*

16: sp\_bacteriop:\*

17: sp\_archaeap:\*

Score Match Length DB ID

17 58.5 266 13 O42143

18 58.5 28.0 2 Q9AL20

19 58.5 28.0 2 Q9XD90

20 57 27.3 2 Q9LT54

21 56.5 27.0 60.8 10 O22678

22 56 26.8 2127 12 O57294

23 56 26.8 2127 13 Q9JH63

24 55.5 26.6 1272 13 Q90924

25 55.5 26.6 1369 13 Q42414

26 55 26.3 130 11 Q9CVF1

27 55 26.3 132 10 Q9XD9

28 55 26.3 132 10 Q9SXCL

29 55 26.3 144 11 Q9D887

30 55 26.3 188 2 Q49387

31 55 26.3 192 2 Q49390

32 55 26.3 261 3 Q9C2U0

33 55 26.3 392 2 Q93CN3

34 55 26.3 439 17 Q8ZwR8

35 55 26.3 790 5 Q20949

36 54.5 26.1 221 5 O62473

37 54.5 26.1 298 2 O57280

38 54.5 26.1 298 2 Q46161

39 54.5 26.1 298 2 Q46162

40 54.5 26.1 298 2 Q93777

41 54.5 26.1 379 2 O85863

42 54 25.8 309 5 002163

43 54 25.8 310 17 Q9Ye06

44 54 25.8 458 10 Q49922

45 54 25.8 464 10 Q9LDR5

## ALIGNMENTS

## RESULT 1

O42143

AC O42143 ; PRELIMINARY; PRT; 266 AA.

ID 042143 ; DT 01-JAN-1998 (TREMBLrel. 05, Created)

DT 01-JAN-1998 (TREMBLrel. 05, Last sequence update)

DT 01-JUN-2001 (TREMBLrel. 17, Last annotation update)

DE Glucagon I precursor [Contains: Glucagon; Glucagon-like peptide 1A (GLP-1A); Glucagon-like peptide 1B (GLP-1B); Glucagon-like peptide 1C (GLP-1C); Glucagon-like peptide 2 (GLP-2)].

DB Xenopus laevis (African clawed frog).

OS Xukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi; Amphibia; Batrachia; Anura; Mesobatrachia; Pipoidea; Pipidae; Xenopoda; Xenopus.

OC NCBI TaxID=8355;

OX NCBI TaxID=8355;

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

## SUMMARIES

Result No.	Score	Match Length	DB ID	Description
1	11.9	56.9	266 13 O42143	O42143 xenopus lae
2	11.1	53.1	219 13 O42144	O42144 xenopus lae
3	10.1	48.3	220 13 Q9BW19	Q9BW19 hoplobatrach
4	90.5	43.3	178 13 Q91971	Q91971 oncohyynchus
5	88	42.1	72 13 Q91409	Q91409 oncohyynchus
6	88	42.1	178 13 Q91189	Q91189 oncohyynchus
7	88	42.1	206 13 Q91410	Q91410 gallus gallus
8	87	41.6	180 6 Q95LG0	Q95LG0 canis familiaris
9	86.5	41.4	62 13 Q9PRW9	Q9PRW9 scylliorhinus
10	86.5	41.4	160 13 Q9PUR1	Q9PUR1 petromyzon
11	83	39.7	121 13 Q9DDB6	Q9DDB6 brachydanius
12	82	39.2	204 13 O12956	O12956 heloderma
13	77	36.8	96 13 Q9PG43	Q9PG43 amblonychites
14	75	35.9	120 13 Q9PUB0	Q9PUB0 petromyzon
15	62	29.7	347 16 Q9XW3	Q9XW3 rhizobium m
16	61	193	5 Q9V712	Q9V712 drosophila

FT SIGNAL

RT "The Xenopus proglucagon gene encodes novel GLP-1-like peptides with insulinotropic properties";

RT Proc. Natl. Acad. Sci. U.S.A. 94: 7915-7920 (1997).

RL -!- FUNCTION: PROMOTES HYDROLYSIS OF GLYCOCGEN AND LIPIDS, AND RAISES THE BLOOD SUGAR LEVEL.

CC -!- ALTERNATIVE PRODUCTS: 2 ISOFORMS; 1 (SHOWN HERE) AND 2; ARE PRODUCED BY ALTERNATIVE SPlicing.

CC -!- SIMILARITY: BELONGS TO THE GLUCAGON FAMILY.

CC DR EMBL; AF004432; AB65660.1; -

CC DR HSSP; P0127; 1GEN

CC DR Intero; IPR00052; Glucagon.

DR PRINS; PR00123; hormone2; 5.

DR SMART; SM00070; GLUCA; 5.

DR PROSITE; PS00260; GLUCAGON; 5.

KW Glucagon family; Hormone; Signal; Cleavage on pair of basic residues; Multisene family; Alternative splicing.

KW Potential?

FT PEPTIDE 53 81 GLUCAGON-LIKE PEPTIDE 1A.  
 FT PEPTIDE 97 133 GLUCAGON-LIKE PEPTIDE 1B.  
 FT PEPTIDE 142 173 GLUCAGON-LIKE PEPTIDE 1C.  
 FT PEPTIDE 180 211 GLUCAGON-LIKE PEPTIDE 2.  
 FT PEPTIDE 227 259 MISSING (IN 1SFORM 2).  
 FT VASPLIC 214 261 Hoplobatrachus rugulosus,  
 SEQUENCE 266 AA; 30951 MW; 544FTBBC20AFAF72C CRC64;  
 NCBI\_TaxID=110072;

Query Match 56.9%; Score 119; DB 13; Length 266;  
 Best Local Similarity 62.5%; Pred. No. 8.9e-08;  
 Matches 20; Conservative 8; Mismatches -4; Indels 0; Gaps 0;

Qy 1 HGGTFTSDLSKQMBEAVRLFIWLNKGPS 32  
 Db 97 HAEGTFTSDVTOQDKEKAKEFIDWLNGPS 128

RESULT 2  
 ID 042144 PRELIMINARY; PRT; 219 AA.  
 AC 042144; 05. Created  
 DT 01-JAN-1998 (TREMBLrel. 05, Last sequence update)  
 DT 01-JUN-2001 (TREMBLrel. 17, Last annotation update)  
 DE Glucagon II precursor [Contains: Glucagon; glucagon-like peptide 1A  
 DE (GLP-1); glucagon-like peptide 1B (GLP-1B); glucagon-like peptide 1C  
 DE (GLP-1C)].  
 OS Xenopus laevis (African clawed frog).  
 OC Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
 OC Amphibia; Batrachia; Anura; Mesobatrachia; Pipoidea; Pipidae;  
 OC Xenopodinae; Xenopus.  
 OX NCBI\_TaxID=8355;

RN [1] 4  
 RP SEQUENCE FROM N.A.  
 RC TISSUE= PANCREAS; PubMed=9223287;  
 RX MEDLINE=97368292; Irwin D.M., Satturajah M., Wen Y., Brubaker P.L., Pederson R.A., Wheeler M.B., RT "The Xenopus proglucagon gene encodes novel GLP-1-like peptides with insulinotropic properties." Proc Natl Acad Sci U S A, 94,7915-7920(1997).  
 RL -I- FUNCTION: PROMOTES HYDROLYSIS OF GLICOGEN AND LIPIDS, AND RAISES THE BLOOD SUGAR LEVEL.  
 CC -I- SIMILARITY: BELONGS TO THE GLUCAGON FAMILY.  
 DR EMBL: AF004433; AAB65661.1; -.  
 DR HSSP; P0174; 1GDN.  
 DR InterPro; IPR00532; Glucagon.  
 DR Pfam; PF00123; hormone 4.  
 DR PRINTS; PR00275; GLUCAGON.  
 DR SMART; SM00070; GLUCA 4.  
 DR PROSITE; PS00260; GLUCAGON.  
 DR SMART; SM00070; GLUCA 4.  
 DR PROSITE; PS00260; GLUCAGON; UNKNOWN 4.  
 SQ SEQUENCE 220 AA; 25615 MW; C7D52657F89E381 CRC64;

RESULT 4  
 ID 091971 PRELIMINARY; PRT; 178 AA.  
 AC 091971; Q91408; Q91188; Q92169;  
 DT 01-NOV-1996 (TREMBLrel. 01, Created)  
 DT 01-NOV-1996 (TREMBLrel. 01, Last sequence update)  
 DT 01-JUN-2001 (TREMBLrel. 17, Last annotation update)  
 DE Glucagon I precursor.  
 OS Oncorhynchus mykiss (Rainbow trout) (Salmo gairdneri).  
 OC Bukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
 OC Actinopterygii; Neopterygii; Teleostei; Euteleostomi;  
 OC Protacanthopterygii; Salmoniformes; Salmonidae; Oncorhynchus.  
 OX NCBI\_TaxID=8022;

RN [1] 4  
 RP SEQUENCE FROM N.A.; AND ALTERNATIVE SPLICING.  
 RC TISSUE=DISTAL SMALL INTESTINE, AND PANCREAS;  
 RX MEDLINE=77295739; PubMed=776376;

Qy 1 HGGTFTSDLSKQMBEAVRLFIWLNKGPS 32  
 Db 97 HAEGTFTSDVTOQDKEKAKEFIDWLNGPS 128

RESULT 3  
 ID Q8WNL9 PRELIMINARY; PRT; 220 AA.  
 AC Q8WNL9;

KW	Glucagon family;	Hormone; Cleavage on pair of basic residues; Signal;
KW	Alternative splicing;	Multigene family;
SIGNAL	1	POTENTIAL?
PEPTIDE	?	49
PEPTIDE	52	80
PEPTIDE	85	120
PEPTIDE	137	169
PEPTIDE	124	178
VARSPIC	178 AA;	20034 MW;
SEQUENCE		5CF6980CF2A9D586 CRC44;
SQ		
Query	Match	Score 43.3%; DB 13; Length 178;
Best	Local Similarity	50.0%; Pred. No. 0.00037;
Matches	19;	Mismatches 5; Indels 1; Gaps
Qy	1 HGEGETFTSDLSKOMEERAVRLFIEWLNGGPGSSGAP 36	
Db	52 HSEGFTFSNDYKSYQERMAQDFTQWLMMN-SKRGAP 86	

RESULT 5

Q91409	Q91409	PRELIMINARY;	PRT;	72 AA.
ID	ID			
AC	Q91409;	Q91232;		
DT	01-NOV-1995	(TREMBLrel).	01	Created
DT	01-NOV-1996	(TREMBLrel).	01	Last sequence update
DT	01-DEC-2001	(TREMBLrel).	19	Last annotation update
PROLG	YACON			
OS	Oncorhynchus tshawytscha	(Chinook salmon)	(King salmon)	
OC	Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Buteleostomi;			
OC	Actinopterygii; Neopterygii; Teleostei; Buteleosteoi;			
OC	Protacanthopterygii; Salmoniformes; Salmonidae; Oncorhynchus.			
NCBI_TAXID	74940;			
RN	[1]			
SEQUENCE FROM N.A.				
MRNA	SEQUENCE FROM N.A.			
MRP	SEQUENCE FROM N.A.			
RX	SEQUENCE FROM N.A.			
RA	SEQUENCE FROM N.A.			
RA	Irwin D.M.; Wong J.;			
RA	"trout and chicken proglucagon: alternative splicing generates mRNA			
RT	transcripts encoding glucagon-like peptide 2.;"			
RT	Mol. Endocrinol. 9:267-277(1995).			
DR	EMBL: S78474; AAD14283.1; -;			
DR	EMBL: U19940; AAC59670.1; -;			
DR	HSSP: P01274; IGCN			
DR	InterPro: IPR00532; Glucagon.			
DR	InterPro: IPR00532; Glucagon.			

Query	Match	Score	DB	Length	72;
PRINTER	PRINTER	42.1%	88;	13;	
SMART	SMART	44.8%	DB	0.00029;	
PROSIE	PROSIE	13;	Matches	10;	Indels
NON TIR	NON TIR	1	Conservative	6;	0;
SEQUENCE	SEQUENCE	72	AA:	8293	Gaps
				MM:	
				8584352B1C260A31	
				CRC64;	

RESULT 6  
29.1.189  
Q91189; Q92168; Q91189; Q92168; PRELIMINARY; PRT; 178 AA.  
AC (TREMBLrel. 01. Created)  
DT 01-NOV-1996 (TREMBLrel. 01. Last sequence update)  
DT 01-JUN-2001 (TREMBLrel. 01. Last annotation update)  
Glucagon II precursor.  
Oncorhynchus mykiss (Rainbow trout) (*Salmo Gairdneri*).  
Bukaria; Metza; Chordata; Craniata; Vertebrata; Buteleostomi;  
Actinopterygii; Neopterygii; Teleostei; Euteleostei;  
Protacanthopterygii; Salmoniformes; Salmonidae; Oncorhynchus.  
NCBI\_TaxID=8022;  
OX

RP SEQUENCE FROM N.A., AND ALTERNATIVE SPLICING.  
 RC TISSUES=DISTAL SMALL INTESTINE, AND PANCREAS;  
 RX MEDLINE=55295739; PubMed=77765976;  
 RA Irwin D.M.; Wong J.;  
 RT "trout and chicken proglucagon: alternative splicing generates mRNA  
 RT transcripts encoding glucagon-like peptide 2";  
 RL Mol. Endocrinol. 9:267-277 (1995).  
 CC -1- FUNCTION: PROMOTES HYDROLYSIS OF GLYCOGEN AND LIPIDS, AND RAISES  
 CC THE BLOOD SUGAR LEVEL (BY SIMILARITY).  
 CC -1- ALTERNATIVE PRODUCTS: 2 ISOFORMS: INTESTINAL (SHOWN HERE) AND  
 CC PANCREATIC; ARE PRODUCED BY ALTERNATIVE SPLICING.  
 CC -1- RESPONSE: PRODUCED IN THE A CELLS OF THE ISLETS OF LANGERHANS IN  
 CC PANCREAS; A DROP IN BLOOD SUGAR CONCENTRATION.  
 CC -1- SIMILARITY: BELONGS TO THE GLUCAGON FAMILY.  
 DR EMBL: U19914; AAC59658.1; -;  
 DR EMBL: U19916; AAC60210.1; -;  
 DR EMBL: U19915; AAC60210.1; JOINED.  
 DR EMBL: U19915; AAC60209.1; -;

DR	HSSP / P01274 / 1GEN.			
DR	IntersPro: IP000332; Glucagon.			
DR	Pfam: PF001122; hormone2_3.			
DR	PRINTS: PR00215; GLUCAGON.			
DR	SMART: SM00070; GLUCAGON.			
DR	PROSITE: PS00260; GLUCAGON; UNKNOWN 2.			
KW	Glucagon family Hormone; Cleavage on pair of basic residues; Signal			
KW	Alternative splicing; Multigene family.			
KW	POTENTIAL.			
FT	SIGNAL, 1?			
FT	PEPTIDE 2	49	GRPP (GLICENTINE RELATED POLYPEPTIDE).	
FT	PEPTIDE 52	80	GLUCAGON.	
FT	PEPTIDE 85	120	GLUCAGON-LIKE PEPTIDE 1.	
FT	PEPTIDE 137	169	GLUCAGON-LIKE PEPTIDE 2.	
FT	VARSPLIC 124	178	MISSING (IN PANCREATIC ISOFORM).	
FT	SEQUENCE 178 AA;	19998 MW;	E89D7D3866CD91C56 CRC64;	
Query Match	Best Local Similarity	42.1%	Score 88;	DB 13;
Qy	Matches 13; Conservative	44.8%	Pred. No. 0.0008;	Length 178;
Db	10; Mismatches 6;	Indels 0;	Gaps	
RESULT 7				
Q91410		PRELIMINARY;	PRT;	206 AA.
Q91410;				
AC				
DT	01-NOV-1996	(TREMBLrel. 01, Created)		
DT	01-NOV-1996	(TREMBLrel. 01, Last sequence update)		
DT	01-DEC-2001	(TREMBLrel. 19, Last annotation update)		
DE	Proglucagon.			
GN	PROGUCAGON.			
OS	Gallus gallus (Chicken).			
OC	Bukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;			
OC	Archosauria; Aves; Neognathae; Galliformes; Phasianidae; Phasianinae			

OX [1] - NCBI - TAXID=9031;  
 RP SEQUENCE FROM N. A.  
 RX MEDLINE=5295739; PubMed=7776976;  
 RA Irwin D.M.; Wong J.;  
 RT "trout and chicken proglucagon: alternative splicing generates mRNA  
 transcripts encoding glucagon-like peptide 2.,";  
 RL Mol. Endocrinol. 9:267-277 (1995).  
 EMBL S78477; AB34506.1;  
 DR P01274; IGCN.  
 DR InterPro:IPR000532; Glucagon.  
 DR PFAM:PF01231; Hormone2; 3.  
 DR PRINTS:PR00275; GLUCAGON.  
 DR SMART:SM00070; GLUCAGON.  
 DR PROSITE:PS00260; GLUCAGON; 3.  
 DR PROTEIN:PS00260; GLUCAGON; 3.  
 DR PROTEIN:PS00260; GLUCAGON; 3.

FT	PEPTIDE	1	29	GLUCAGON-29.
FT	PEPTIDE	1	33	GLUCAGON-33.
FT	NON_CONS	33	34	
FT	PEPTIDE	34	62	GLUCAGON-LIKE PEPTIDE.
SQ	SEQUENCE	62 AA;	2720 MW;	CSFP487C12059CD1 CRC64;
Query Match	41.4%;	Score 86.5;	DB 13;	Length 62;
Best Local Similarity	45.9%;	Pred. No. 0.00038;		
Matches	17;	Conservative	5;	Mismatches 12;
				Indels 3; Gaps
Qy	1 HSEGTTFTSDSKQMEEAAVRLFIBWL---RNKGPSGG 34			
Db	1 HSEGTTFTSDSKYKLENQQAQFVWLMSTKENGHARG 37			
RESULT 10				
Q9PDR1	PRELIMINARY;	PRT;	160 AA.	
ID	Q9PDR1; Q9PZ28; Q9PZ27;			
AC	O9PDR1;			
DT	01-MAY-2000 (TRMBLrel. 13; Created)			
DT	01-MAY-2000 (TRMBLrel. 13; Last sequence update)			
DT	01-DEC-2001 (TRMBLrel. 19; Last annotation update)			
DE	Glucagon I precursor [Contains: Glucagon; Glucagon-like peptide 1 (GLP-1); Glucagon-like peptide 2 (GLP-2)].			
OS	Petromyzon marinus (Sea Lamprey).			
OC	Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Hyperoartia; Petromyzontiformes; Petromyzontidae; Petromyzon.			
NCBI_TaxID	7757;			
RN	[1]			
RP	SEQUENCE FROM N.A.			
RC	TISSUE=INTESTINE;			
RX	MEDLINE=20022986; PubMed=10555286;			
RA	Irwin D.M., Huner O., Youson J.H.;			
RT	"Lamprey proglucagon and the origin of glucagon-like peptides.";			
RL	Mol. Biol. Evol. 16:1548-1557(1999).			
RN	[2]			
RP	SEQUENCE OF 43-71 AND 82-113.			
RC	TISSUE=INTESTINE;			
RX	MEDLINE=94010172; PubMed=8405897;			
RA	Conlon J.M., Nielsen P.F., Youson J.H.;			
RT	"Primary structures of glucagon and glucagon-like peptide isolated from the intestine of the parasitic phase lamprey Petromyzon marinus.";			
RT	Gen. Comp. Endocrinol. 91:96-104(1993).			
CC	-I- FUNCTION: PROMOTES HYDROLYSIS OF GLYCOGEN AND LIPIDS, AND RAISES THE BLOOD SUGAR LEVEL.			
CC	-I- SIMILARITY: BELONGS TO THE GLUCAGON FAMILY.			
DR	EMLB; AF159707; AAP09186.1; -.			
DR	HSSP: P01275; 1B00.			
DR	InterPro: IPR000532; Glucagon.			
DR	PFAM: PF0123; hormone; 2.			
DR	PRINTS: PR00275; GLUCAGON.			
DR	SMART: SM00070; GLUCA; 2.			
DR	PROSITE: PS00260; GLUCAGON; 2.			
RW	Glucagon family; Hormone; Signal; Cleavage on pair of basic residues; Multigene family.			
RW	FT SIGNAL 1 22 POTENTIAL.			
FT	PEPTIDE 43 71 GLUCAGON.			
FT	PEPTIDE 82 113 GLUCAGON-LIKE PEPTIDE 1.			
FT	PEPTIDE 130 160 GLUCAGON-LIKE PEPTIDE 2.			
SQ	SEQUENCE 160 AA; 18042 MW; 9A52C330D5A74072 CRC64;			
Query Match	41.4%;	Score 86.5;	DB 13;	Length 160;
Best Local Similarity	51.5%;	Pred. No. 0.0011;		
Matches	17;	Conservative	4;	Mismatches 9; Indels 3; Gaps
Qy	1 HSEGTTFTSDLSKQMEEAAVRLFIBWL---RNKG 30			
Db	43 HSEGTTFTSDSKYKLENQQAQFVWLMSTKENGHARG 75			

ID	Q9DDE6	PRELIMINARY;	PRT;	121 AA.	DR	EMBL; U77611; AAB511128.1; -.
AC	Q9DDE6	Created)			DR	HSSP; P01274; IGCN.
DT	01-MAR-2001	(TREMBLrel. 16, Last sequence update)			DR	InterPro; IPR000332; Glucagon.
DT	01-MAR-2001	(TREMBLrel. 16, Last annotation update)			DR	PFam; PF00122; hormone2; 3.
DT	01-DEC-2001	(TREMBLrel. 19, Last annotation update)			DR	PRINTS; PR00375; GLUCAGON.
DE	Glucagon, polyprotein.				DR	SMART; SM00070; GLUC; 3.
CN	GGC OR GLU.				DR	PROSITE; PS00260; GLUCAGON; 2.
OS	Brachydanio rerio (Zebrafish) (Zebra daniel).				RW	Glucagon family; Hormone; Cleavage on pair of basic residues; Signal;
OC	Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Ostariophysi; Cypriniformes;				RW	Alternative splicing.
OC	Actinopterygii; Neopterygii; Teleostei; Ostariophysi; Cypriniformes;				RW	Signal.
OC	Cyprinidae; Danio.				FT	SIGNAL; 1 20
OC	NCBI_TaxID=7955;				FT	PEPTIDE 21 50
OX					FT	GLUCAGON
RN	[1]				FT	GLUCAGON-LIKE PEPTIDE 1.
RP	SEQUENCE FROM N.A.				FT	GLUCAGON-LIKE PEPTIDE 2.
RX	Medline=994251930; PubMed=10495291;				FT	D -> E (IN ISOFORM LPI).
RA	Argentan P., Zucchin B., Bortolussi M.;				FT	MISSING (IN ISOFORM LPI).
RT	"Early appearance of pancreatic hormone-expressing cells in the zebrafish embryo."				FT	VARSPLIC 150 204 MISSING (IN ISOFORM LPI).
RT	Mech. Dev. 87:217-221(1999).				FT	SEQUENCE 204 AA; 23553 MW; B132E5F46873E72 CRC64;
RL	ENB; AJ13697; CAC20108.1; -.				Query Match	Score 39.2%; DB 13; Length 204;
DR	HSSP; P01274; IGCN				Best Local Similarity	48.3%; Pred. No. 0.0059;
DR	ZFIN; ZDB-GENE-010129-1; geg.				Matches	6; Mismatches 9; Indels 0; Gaps 0;
DR	InterPro; IPR00532; Glucagon.				Qy	1 HGEGETFTSDLSKQMEEEAVRLFIEWLKNGP 29
DR	PFam; PF00123; hormone2; 2.				Db	116 HADGRYDISSYLEGQAKEYFLAWLNG 144
DR	PRINTS; PS00275; GLUCAGON.					
DR	SMART; SM00070; GLUC; 2.					
DR	PROSITE; PS00260; GLUCAGON; 1.					
KW	Polyprotein.					
FT	CHAIN 49 79					
FT	CHAIN 88 121 AA; 13537 MW; A85385F690DA180P CRC64;					
SQ						
Query Match	Score 39.7%; DB 13; Length 121;				RESULT 13	Q9PG43
Best Local Similarity	45.2%; Pred. No. 0.0024;				ID	Q9DG43
Matches	9; Mismatches 8; Indels 0; Gaps 0;				PRELIMINARY;	PRT; 96 AA.
Qy	1 HGEGETFTSDLSKQMEEEAVRLFIEWLKNGP 31				AC	Q9DG43;
Db	88 HAEGBTYTSDVSSYLQDQAQRVARLKGQP 118				DT	01-MAR-2001 (TREMBLrel. 16, Created)
					DT	01-MAR-2001 (TREMBLrel. 16, Last sequence update)
					DR	01-DEC-2001 (TREMBLrel. 19, Last annotation update)
					DS	Proglucagon (Fragment).
					OS	Ambloplites rupestris.
					OC	Actinopterygii; Neopterygii; Teleostei; Vertebrates; Euteleostomi;
					OC	Acanthomorpha; Acanthopterygii; Neoteleostei;
					OC	Centrarchidae; Ambloplites.
					OX	NCBI_TaxID=109273;
					RN	[1]
					RP	SEQUENCE FROM N.A.
					RA	Al-Mahrouki A.A., Irwin D.M., Youson J.H.;
					RT	"Rock Bass Proglucagon."
					RL	Submitted (SEP-1999) to the EMBL/GenBank/DBJ databases.
					DR	EMBL; AF19049; AAC16778.1; -.
					DR	HSSP; P01274; IGCN.
					DR	InterPro; IP000332; Glucagon.
					DR	PFam; PF00122; hormone2; 2.
					DR	PRINTS; PR00215; GLUCAGON.
					DR	SMART; SM00070; GLUC; 2.
					DR	PROSITE; PS00260; GLUCAGON; UNKNOWN_1.
					FT	NON-TER 1 1
					FT	CHAIN 1 >29
					FT	GLUCAGON
					FT	GLUCAGON-LIKE PEPTIDE 1.
					FT	GLUCAGON-LIKE PEPTIDE 2.
					FT	NON-TER 96 96
					FT	>96 >96
					FT	SEQUENCE 96 AA; 11225 MW; 6435033BBDG00CE CRC64;
					Query Match	Score 36.8%; DB 13; Length 96;
					Best Local Similarity	40.0%; Pred. No. 0.012;
					Matches	6; Mismatches 15; Indels 0; Gaps 0;
Qy	1 HGEGETFTSDLSKQMEEEAVRLFIEWLKNGPSSGA 35				Qy	1 HGEGETFTSDLSKQMEEEAVRLFIEWLKNGPSSGA 35
Db	1 HSQGFTFTNDYTYLLEDQAOQDFIRMLNNKRNKSGAA 35				Db	1 HSQGFTFTNDYTYLLEDQAOQDFIRMLNNKRNKSGAA 35
					RESULT 14	
					O9PURO	
					ID	O9PURO
					AC	Q9PURO;

01-MAY-2000 (TREMBLrel. 13; Created)  
 DT 01-MAY-2000 (TREMBLrel. 13; Last sequence update)  
 DT 01-DEC-2001 (TREMBLrel. 19; Last annotation update)  
 DE Glucagon II precursor (Contains: Glucagon; Glucagon-like peptide (GIP).)  
 DE  
 IS Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Hyperoartia;  
 OC Percomyzontiformes; Petromyzontidae; Petromyzon.  
 OC NCBITaxID=7557;  
 OX [1]  
 RN  
 SEQUENCE FROM N.A.  
 RC TISSUE=INTESTINE;  
 RX MEDLINE=20022986; PubMed=10555286;  
 RA Irwin D.M., Hunter O., Youson J.H.;  
 RT "Lamprey proglucagon and the origin of glucagon-like peptides.";  
 RL Mol. Biol. Evol. 16:1548-1557 (1999).  
 CC -1- FUNCTION: PROMOTES HYDROLYSIS OF GLYCOGEN AND LIPIDS, AND RAISES THE BLOOD SUGAR LEVEL.  
 CC -1- SIMILARITY: BELONGS TO THE GLUCAGON FAMILY.  
 DR EMBL; AF159708; AAF09187.1; -.  
 DR HSSP; P01275; 1EHO  
 DR InterPro; IPR00532; Glucagon.  
 DR Pfam; PF00123; Hormone\_2.  
 DR PRINTS; PRO0275; GLUCAGON.  
 DR SMART; SM00070; GLUCA\_2.  
 DR PROSITE; PS00266; GLUCAGON\_2.  
 DR Glucagon family; Hormone; Signal; Cleavage on pair of basic residues;  
 KW Multigene family.  
 PT SIGNAL 1 ? POTENTIAL.  
 PT PEPTIDE 44 72 GLUCAGON-LIKE PEPTIDE.  
 PT PEPTIDE 89 120 MW: FBDBE67B36E198D8 CRC64;  
 SQ SEQUENCE 120 AA; 13397 MW:  
 Query Match 35.9%; Score 75; DB 13; Length 120;  
 Best Local Similarity 36.7%; Pred. No. 0.028;  
 Matches 11; Conservative 10; Mismatches 9; Indels 0; Gaps 0;  
 OX 1 HGEHTFTSDLRQMEEAIVRLFIEWLNGG 30  
 DB 89 HSDGSFINDMMVYMLDRMSAKNFWLKGQ 118

RESULT 15  
 Q92XW3 PRELIMINARY; PRT; 347 AA.  
 ID Q92XW3  
 AC  
 DT 01-DEC-2001 (TREMBLrel. 19; Created)  
 DT 01-DEC-2001 (TREMBLrel. 19; Last sequence update)  
 DT 01-DEC-2002 (TREMBLrel. 20; Last annotation update)  
 DE Hypothetical Protein RA1127.  
 GN RA1127 OR SWA2063  
 OS Rhizobium meliloti (Sinorhizobium meliloti).  
 OG Plasmid psymA (megaplasmid 1).  
 OC Rhizobiaceae; Sinorhizobium.  
 OX NCBITaxID=362;  
 RN SEQUENCE FROM N.A.  
 RC STRAIN=1021;  
 RX MEDLINE=139509; PubMed=1141432;  
 RA Barnett M.J., Fisher R.F., Jones T., Komp C., Abola A.P.,  
 RA Barley-Hubler F., Bowser L., Capela D., Gallibert F., Gonzy J.,  
 RA Gurjai M., Hong A., Huijzer L., Hyman R.W., Kahn D., Kahn M.J.,  
 RA Kalman S., Keating D.H., Palm C., Peck M.C., Surycki R., Weil D.H.,  
 RA Yeh K.-C., Davis R.W., Federici N.A., Long S.R.;  
 RT "Nucleotide sequence and predicted function of the entire  
 RT Sinorhizobium meliloti psymA megaplasmid.";  
 RL Proc. Natl. Acad. Sci. U.S.A. 98:9883-9888 (2001).  
 DR E007238; AAK65785 1;  
 DR InterPro; IPR001005; Myb-DNA binding.  
 DR PROSITE; PS00037; MYB\_1; UNKNOWN.  
 KW Hypothetical Protein; Plasmid; Complete proteome.  
 SQ 347 AA; 38706 MW; 2BDBEF2867AD0475C CRC64;